10/537102 JC09 Rec'd PCT/PTO 02 JUN 2005 PCT/FI2003/000949

## WO 2004/055055

## SEQUENCE LISTING

<1	10> 3	Lice	ntia	Ltd												
<12	20> 1	Nove:	l pro	oteir	n and	l use	s tl	iered	of							
<13	30> 4	1055	2				•									
			0/433 -12-1		L											
<16	50> 3	32				•										
<17	70> I	Pater	ntIn	Ver.	2.1											
<21 <21	.0> 1 .1> 1 .2> I .3> F	.479 NA	sapi	.ens												
<22 <22	0> 1> C	DS						-								
<22	2> (	1)	(147	9)		•						,				
atg	HIS	ccc	cac His	cgt Arg	Asp	ccg Pro	aga Arg	ggc	ctc Leu 10	Trp	ctc Leu	ctg Leu	ctg Leu	ccg Pro 15	tcc Ser	48
ttg Leu	tcc Ser	ctg Leu	ctg Leu 20	Leu	ttt Phe	gag Glu	gtg Val	gcc Ala 25	Arg	gct Ala	ggc	cga Arg	gcc Ala 30	gtg Val	gtt Val	96
agc Ser	tgt Cys	cct Pro 35	Ala	gcc Ala	tgc Cys	ttg Leu	tgc Cys 40	Ala	agc Ser	aac Asn	atc Ile	ctc Leu 45	agc Ser	tgc Cys	tcc Ser	144
aag Lys	cag Gln 50	cag Gln	ctg Leu	ccc Pro	aat Asn	gtg Val 55	ccc	cat His	tcc Ser	ttg Leu	ccc Pro 60	agt Ser	tac Tyr	aca Thr	gca Ala	192
cta Leu 65	ctg Leu	gac Asp	ctc Leu	agt Ser	cac His 70	aac Asn	aac Asn	ctg Leu	agc Ser	cgc Arg 75	ctg Leu	cgg Arg	gcc Ala	gag Glu	tgg Trp 80	240
acc Thr	ccc Pro	acg Thr	cgc Arg	ctg Leu 85	acc Thr	caa Gln	ctg Leu	cac His	tcc Ser 90	ctg Leu	ctg Leu	ctg Leu	agc Ser	cac His 95	aac Asn	288
cac His	ctg Leu	aac Asn	ttc Phe 100	atc Ile	tcc Ser	tct Ser	gag Glu	gcc Ala 105	ttt Phe	tcc Ser	ccg Pro	gta Val	ccc Pro 110	aac Asn	ctg Leu	336
cgc Arg	tac Tyr	ctg Leu 115	gac Asp	ctc Leu	tcc Ser	tcc Ser	aac Asn 120	cag Gln	ctg Leu	cgt Arg	aca Thr	ctg Leu 125	gat Asp	gag Glu	ttc Phe	384
ctg Leu	ttc Phe 130	agt Ser	gac Asp	ctg Leu	caa Gln	gta Val 135	ctg Leu	gag Glu	gtg Val	ctg Leu	ctg Leu 140	ctc Leu	tac Tyr	aat. Asn	aaç Asn	432

														•		
cac His 145	: Ile	atg Met	gcg	gtg Val	gac Asp 150	Arg	tgc Cys	gcc Ala	ttc Phe	gat Asp 155	Asp	atg Met	gcc	cag Gln	ctg Leu 160	480
caç Glr	aaa Lys	ctc Leu	tac Tyr	ttg Leu 165	Ser	cag Gln	aac Asn	cag Gln	ato Ile 170	Ser	cgc	ttc Phe	cct	ctg Leu 175	Glu	528
ct <u>c</u> Lev	gtc Val	aag Lys	gaa Glu 180	Gly	gcc Ala	aag Lys	cta Leu	ccc Pro 185	Lys	cta Leu	acg Thr	ctc Leu	ctg Leu 190	Āsp	ctc Leu	576
tct Ser	tct Ser	aac Asn 195	Lys	ctg Leu	aag Lys	aac Asn	ttg Leu 200	cca Pro	ttg Leu	cct Pro	gac Asp	ctg Leu 205	cag Gln	aag Lys	ctg Leu	624
ecg Pro	gcc Ala 210	Trp	atc Ile	aag Lys	aat Asn	999 Gly 215	ctg Leu	tac Tyr	cta Leu	cat His	aac Asn 220	aac Asn	ccc Pro	ctg Leu	aac Asn	672
tgo Cys 225	gac Asp	tgt Cys	gag Glu	ctc Leu	tac Tyr 230	Gln	ctg Leu	ttt Phe	tca Ser	cac His 235	Trp	cag Gln	tat Tyr	cgg Arg	cag Gln 240	720
ctg Leu	agc Ser	tcc Ser	gtg Val	atg Met 245	gac Asp	ttt Phe	caa Gln	gag Glu	gat Asp 250	ctg Leu	tac Tyr	tgc Cys	atg Met	aac Asn 255	tcc Ser	768
aag Lys	aag Lys	ctg Leu	cac His 260	aat Asn	gtc Val	ttc Phe	aac Asn	ctg Leu 265	agt Ser	ttc Phe	ctc Leu	aac Asn	tgt Cys 270	Gly	gag Glu	816
tac Tyr	aag Lys	gag Glu 275	cgt Arg	.gcc Ala	tgg Trp	gag Glu	gcc Ala 280	cac His	ctg Leu	ggt Gly	gac Asp	acc Thr 285	ttg Leu	atc Ile	atc Ile	864
aag Lys	tgt Cys 290	gac Asp	acc Thr	aag Lys	cag Gln	caa Gln 295	gj <sup>λ</sup> aaa	atg Met	acc Thr	aag Lys	gtg Val 300	tgg Trp	gtg Val	aca Thr	cca Pro	912
	aat Asn															960
tct Ser	aag Lys	gat Asp	gly ggc	agt Ser 325	ctt Leu	ctt Leu	ttc Phe	cag Gln	cag Gln 330	gtg Val	cag Gln	gtc Val	gag Glu	gac Asp 335	ggt Gly	1008
ggt Gly	gtg Val	tat Tyr	acc Thr 340	tgc Cys	tat Tyr	gcc Ala	atg Met	gga Gly 345	gag Glu	act Thr	ttc Phe	aat Asn	gag Glu 350	aca Thr	ctg Leu	1056
tct Ser	gtg Val	gaa Glu 355	ttg Leu	aaa Lys	gtg Val	cac His	aat Asn 360	ttc Phe	acc Thr	ttg Leu	cac His	gga Gly 365	cac His	cat His	gac Asp	1104
Thr	ctc Leu 370	Asn	Thr	Ala	Tyr	Thr 375	Thr	Leu	Val	Gly	380 280	Ile	Leu	Ser	Val	1152
gtc	ctg	gtc	ctc	ata	tac	cta	tac	ctc	acc	cct	tgc	cgc	tgc	tgg	tgc	1200

Val 385	Leu	Val	Leu	Ile	Tyr 390		Tyr	Leu	Thr	Pro 395	_	Arg	Cys	Trp	Cys 400	
cgg Arg	ggt Gly	gta Val	gag Glu	aag Lys 405	Pro	tcc Ser	agc Ser	cat His	caa Gln 410	gga Gly	gac Asp	agc Ser	ctc Leu	agc Ser 415	tct Ser	1248
tcc Ser	atg Met	ctt Leu	agt Ser 420	acc Thr	aca Thr	ccc Pro	aac Asn	cat His 425	gat Asp	cct Pro	atg Met	gct Ala	ggt Gly 430	Gly 999	gac Asp	1296
aaa Lys	gat Asp	gat Asp 435	ggt Gly	ttt Phe	gac Asp	cgg Arg	cgg Arg 440	gtg Val	gct Ala	ttc Phe	ctg Leu	gaa Glu 445	cct Pro	gct Ala	gga Gly	1344
cct Pro	999 Gly 450	cag Gln	ggt Gly	caa Gln	aac Asn	ggc Gly 455	aag Lys	ctc Leu	aag Lys	cca Pro	ggc Gly 460	aac Asn	acc Thr	ctg Leu	cca Pro	1392
gtg Val 465	cct Pro	gag Glu	gcc Ala	aca Thr	ggc Gly 470	aag Lys	ggc	caa Gln	cgg Arg	agg Arg 475	atg Met	tcg Ser	gat Asp	cca Pro	gaa Glu 480	1440
						tct Ser						gtg Val				1479
<212	> 49 2> PF	TS	sapie	ens							,					
<400 Met 1		Pro	His	Arg 5	Asp	Pro	Arg	Gly	Leu 10	Trp	Leu	Leu	Leu	Pro 15	Ser	
Leu	Ser	Leu	Leu 20	Leu	Phe	Glu	Val	Ala 25	Arg	Ala	Gly	Arg	Ala 30	Val	Val	
Ser	Cys	Pro 35	Ala	Ala	Cys	Leu	Cys 40	Ala	Ser	Asn	Ile	Leu 45	Ser	Cys	Ser	
Lys	Gln 50	Gln	Leu	Pro	Asn	Val 55	Pro	His	Ser	Leu	Pro 60	Ser	Tyr	Thr	Ala	
Leu 65	Leu	Asp	Leu	Ser	His 70	Asn	Asn	Leu	Ser	Arg 75	Leu	Arg	Ala	Glu	Trp 80	
Thr	Pro	Thr	Arg	Leu 85	Thr	Gln	Leu	His	Ser 90	Leu	Leu	Leu	Ser	His 95	Asn	
His	Leu	Asn	Phe 100	Ile	Ser	Ser	Glu	Ala 105	Phe	Ser	Pro	Val	Pro 110	Asn	Leu	
Arg		Leu 115	Asp	Leu	Ser	Ser	Asn 120	Gln	Leu	Arg	Thr	Leu 125	Asp	Glu	Phe	
	Phe 130	Ser	Asp	Leu	Gln	Val 135	Leu	Glu	Val	Leu	Leu 140	Leu	Tyr	Asn	Asn	

His 145	Ile	Met	Ala	Val	Asp 150	Arg	Cys	Ala	Phe	Asp 155		Met	Ala	Gln	Leu 160
Gln	Lys	Leu	Tyr	Leu 165		Gln	Asn	Gln	. Ile 170		Arg	Phe	Pro	Leu 175	Glu
Leu	Val	Lys	Glu 180	Gly	Ala	Lys	Leu	Pro 185		Leu	Thr	Leu	Leu 190		Leu
Ser	Ser	Asn 195		Leu	Lys	Asn	Leu 200	Pro	Leu	Pro	Asp	Leu 205		Lys	Leu
Pro	Ala 210		Ile	Lys	Asn	Gly 215	Leu	Tyr	Leu	His	Asn 220	Asn	Pro	Leu	Asn
Суs 225	Asp	Cys	Glu	Leu	Tyr 230	Gln	Leu	Phe	Ser	His 235		Gln	Tyr	Arg	Gln 240
Leu	Ser	Ser	Val	Met 245	Asp	Phe	Gln	Glu	Asp 250		Tyr	Cys	Met	Asn 255	Ser
Lys	· Lys	Leu	His 260	Asn	Val	Phe	Asn	Leu 265	Ser	Phe	Leu	Asn	Cys 270	Gly	Glu
Tyr	Lys	Glu 275	Arg	Ala	Trp	Glu	Ala 280	His	Leu	Gly	Asp	Thr 285		Ile	Ile
Lys	Cys 290	Asp	Thr	Lys	Gln	Gln 295	Gly	Met	Thr	Lys	Val 300	Trp	Val	Thr	Pro
Ser 305	Asn	Glu	Arg	Val	Leu 310	Asp	Glu	Val	Thr	Asn 315	Gly	Thr	Val	Ser	Val 320
Ser	Lys	Asp	Gly	Ser 325	Leu	Leu	Phe	Gln	Gln 330	Val	Gln	Val	Glu	Asp 335	Gly
			340					345					350		Leu
		355					360					365			Asp
	370					375					Cys 380				
385					390					395	Cys				400
				405					410		Asp			415	
Ser	Met	Leu	Ser 420	Thr	Thr	Pro	Asn	His 425	Asp	Pro	Met	Ala	Gly 430	Gly	Asp
		435					440				Leu	445			
	450					455					Gly 460				
Val	Pro	Glu	Ala	Thr	Gly	Lys	Gly	Gln	Arq	Ara	Met	Ser	Asp	Pro	Glu

465 470 475 480

Ser Val Ser Ser Val Phe Ser Asp Thr Pro Ile Val Val 485

<210> 3 <211> 1566 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (1)..(1566) atg tcg tta cgt gta cac act ctg ccc acc ctg ctt gga gcc gtc gtc 48 Met Ser Leu Arg Val His Thr Leu Pro Thr Leu Leu Gly Ala Val Val aga ccg ggc tgc agg gag ctg ctg tgt ttg ctg atg atc aca gtg act Arg Pro Gly Cys Arg Glu Leu Leu Cys Leu Leu Met Ile Thr Val. Thr gtg ggc cet ggt gcc tet ggg gtg tgc ccc acc gct tgc atc tgt gcc 144 Val Gly Pro Gly Ala Ser Gly Val Cys Pro Thr Ala Cys Ile Cys Ala act gac atc gtc agc tgc acc aac aaa aac ctg tcc aag gtg cct ggg 192 Thr Asp Ile Val Ser Cys Thr Asn Lys Asn Leu Ser Lys Val Pro Gly aac ctt ttc aga ctg att aag aga ctg gac ctg agt tat aac aga att 240 Asn Leu Phe Arg Leu Ile Lys Arg Leu Asp Leu Ser Tyr Asn Arg Ile ggg ctt ctg gat tct gag tgg att cca gta tcg ttt gca aag ctg aac 288 Gly Leu Leu Asp Ser Glu Trp Ile Pro Val Ser Phe Ala Lys Leu Asn 85 acc cta att ctt cgt cat aac aac atc acc agc att tcc acg ggc agt 336 Thr Leu Ile Leu Arg His Asn Asn Ile Thr Ser Ile Ser Thr Gly Ser 105 ttt tcc aca act cca aat ttg aag tgt ctt gac tta tcg tcc aat aag 384 Phe Ser Thr Thr Pro Asn Leu Lys Cys Leu Asp Leu Ser Ser Asn Lys ctg aag acg gtg aaa aat gct gta ttc caa gag ttg aag gtt ctg gaa 432 Leu Lys Thr Val Lys Asn Ala Val Phe Gln Glu Leu Lys Val Leu Glu 130 135 gtg ctt ctg ctt tac aac aat cac ata tcc tat ctc gat cct tca gcg 480 Val Leu Leu Tyr Asn Asn His Ile Ser Tyr Leu Asp Pro Ser Ala 150 ttt gga ggg ctc tcc cag ttg cag aaa ctc tac tta agt gga aat ttt Phe Gly Gly Leu Ser Gln Leu Gln Lys Leu Tyr Leu Ser Gly Asn Phe

170

				ccg Pro												576
				tta Leu												624
				aat Asn	Leu											672
				cca Pro												720
-		-		cgt Arg 245				_			_	_		_		768
				cgc Arg												816
ctt Leu	ctg Leu	ctc Leu 275	cag Gln	gat Asp	agc Ser	ttt Phe	atg Met 280	aat Asn	tgc Cys	tct Ser	gac Asp	agc Ser 285	atc Ile	atc Ile	aat Asn	864
ggt	tcc Ser 290	ttt Phe	cgt Arg	gcg Ala	ctt Leu	ggc Gly 295	ttt Phe	att Ile	cat His	gag Glu	gct Ala 300	cag Gln	gtc Val	gjå aaa	gaa Glu	912
				cac His												960
				ggt Gly 325												1008
_	_			tac Tyr						_	_	_		_	_	1056
				gat Asp												1104
				aat Asn												1152
				aga Arg												1200
acc Thr	act Thr	ctt Leu	gct Ala	gct Ala 405	tgc Cys	gtg Val	gcc Ala	agt Ser	atc Ile 410	gtt Val	ttg Leu	gta Val	ctt Leu	ttg Leu 415	tac Tyr	1248
ctc	tat	ctg	act	cca	tgc	ccc	tgc	aag	tgt	aaa	acc	aag	aga	cag	aaa	1296

Leu Tyr Leu Thr Pro Cys Pro Cys Lys Cys Lys Thr Lys Arg Gln Lys 425 aat atg cta cac caa agc aat gcc cat tca tcg att ctc agt cct ggc 1344 Asn Met Leu His Gln Ser Asn Ala His Ser Ser Ile Leu Ser Pro Gly 1392 ccc gct agt gat gcc tcc gct gat gaa cgg aag gca ggt gca ggt aaa Pro Ala Ser Asp Ala Ser Ala Asp Glu Arg Lys Ala Gly Ala Gly Lys aga gtg gtg ttt ttg gaa ccc ctg aag gat act gca gca ggg cag aac 1440 Arg Val Val Phe Leu Glu Pro Leu Lys Asp Thr Ala Ala Gly Gln Asn ggg aaa gtc agg ctc ttt ccc agc gag gca gtg ata gct gag ggc atc Gly Lys Val Arg Leu Phe Pro Ser Glu Ala Val Ile Ala Glu Gly Ile 485 490 cta aag too acg agg ggg aaa tot gac toa gat toa gto aat toa gtg 1536 Leu Lys Ser Thr Arg Gly Lys Ser Asp Ser Asp Ser Val Asn Ser Val 500 505 1566 ttt tct gac aca cct ttt gtg gcg tcc act Phe Ser Asp Thr Pro Phe Val Ala Ser Thr 520 515 <210> 4 <211> 522 <212> PRT <213> Homo sapiens <400> 4 Met Ser Leu Arg Val His Thr Leu Pro Thr Leu Leu Gly Ala Val Val Arg Pro Gly Cys Arg Glu Leu Leu Cys Leu Leu Met Ile Thr Val Thr Val Gly Pro Gly Ala Ser Gly Val Cys Pro Thr Ala Cys Ile Cys Ala Thr Asp Ile Val Ser Cys Thr Asn Lys Asn Leu Ser Lys Val Pro Gly Asn Leu Phe Arg Leu Ile Lys Arg Leu Asp Leu Ser Tyr Asn Arg Ile Gly Leu Leu Asp Ser Glu Trp Ile Pro Val Ser Phe Ala Lys Leu Asn Thr Leu Ile Leu Arg His Asn Asn Ile Thr Ser Ile Ser Thr Gly Ser 100

1

Phe Ser Thr Thr Pro Asn Leu Lys Cys Leu Asp Leu Ser Ser Asn Lys
115 120 125

Leu Lys Thr Val Lys Asn Ala Val Phe Gln Glu Leu Lys Val Leu Glu

135

Val Leu Leu Leu Tyr Asn Asn His Ile Ser Tyr Leu Asp Pro Ser Ala 150 145 Phe Gly Gly Leu Ser Gln Leu Gln Lys Leu Tyr Leu Ser Gly Asn Phe Leu Thr Gln Phe Pro Met Asp Leu Tyr Val Gly Arg Phe Lys Leu Ala Glu Leu Met Phe Leu Asp Val Ser Tyr Asn Arg Ile Pro Ser Met Pro 200 Met His His Ile Asn Leu Val Pro Gly Lys Gln Leu Arg Gly Ile Tyr Leu His Gly Asn Pro Phe Val Cys Asp Cys Ser Leu Tyr Ser Leu Leu 235 Val Phe Trp Tyr Arg Arg His Phe Ser Ser Val Met Asp Phe Lys Asn 245 Asp Tyr Thr Cys Arg Leu Trp Ser Asp Ser Arg His Ser Arg Gln Val 265 Leu Leu Gln Asp Ser Phe Met Asn Cys Ser Asp Ser Ile Ile Asn Gly Ser Phe Arg Ala Leu Gly Phe Ile His Glu Ala Gln Val Gly Glu 295 Arg Leu Met Val His Cys Asp Ser Lys Thr Gly Asn Ala Asn Thr Asp 315 Phe Ile Trp Val Gly Pro Asp Asn Arg Leu Leu Glu Pro Asp Lys Glu 330 Met Glu Asn Phe Tyr Val Phe His Asn Gly Ser Leu Val Ile Glu Ser Pro Arg Phe Glu Asp Ala Gly Val Tyr Ser Cys Ile Ala Met Asn Lys Gln Arg Leu Leu Asn Glu Thr Val Asp Val Thr Ile Asn Val Ser Asn 375 Phe Thr Val Ser Arg Ser His Ala His Glu Ala Phe Asn Thr Ala Phe Thr Thr Leu Ala Ala Cys Val Ala Ser Ile Val Leu Val Leu Leu Tyr Leu Tyr Leu Thr Pro Cys Pro Cys Lys Cys Lys Thr Lys Arg Gln Lys 420 Asn Met Leu His Gln Ser Asn Ala His Ser Ser Ile Leu Ser Pro Gly 440 Pro Ala Ser Asp Ala Ser Ala Asp Glu Arg Lys Ala Gly Ala Gly Lys 455 450 Arg Val Val Phe Leu Glu Pro Leu Lys Asp Thr Ala Ala Gly Gln Asn

465 470 475 480 Gly Lys Val Arg Leu Phe Pro Ser Glu Ala Val Ile Ala Glu Gly Ile 490 Leu Lys Ser Thr Arg Gly Lys Ser Asp Ser Asp Ser Val Asn Ser Val 505 Phe Ser Asp Thr Pro Phe Val Ala Ser Thr 515 <210> 5 <211> 1512 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (1)..(1512) atg acc tgg ttg gtg ctg ctg ggg aca ctg ctc tgc atg ctg cgc gtt 48 Met Thr Trp Leu Val Leu Leu Gly Thr Leu Leu Cys Met Leu Arg Val ggg tta ggc acc ccg gac tcc gag ggt ttc ccg ccc cgt gcg ctc cac Gly Leu Gly Thr Pro Asp Ser Glu Gly Phe Pro Pro Arg Ala Leu His aac tgc ccc tac aaa tgt atc tgc gct gcc gac ctg cta agc tgc act 144 Asn Cys Pro Tyr Lys Cys Ile Cys Ala Ala Asp Leu Leu Ser Cys Thr ggc cta ggg ctg cag gac gtg cca gcc gag tta cct gcc gct act gcg 192 Gly Leu Gly Leu Gln Asp Val Pro Ala Glu Leu Pro Ala Ala Thr Ala gac etc gac etg age cac aac geg etc eag ege etg ege ecc gge tgg 240 Asp Leu Asp Leu Ser His Asn Ala Leu Gln Arg Leu Arg Pro Gly Trp ttg gcg ccc ctc ttc cag ctg cgc gcc ctg cac cta gac cac aac gaa 288 Leu Ala Pro Leu Phe Gln Leu Arg Ala Leu His Leu Asp His Asn Glu cta gat gcg ctg ggt cgc ggc gtc ttc gtc aac gcc agc ggc ctg agg Leu Asp Ala Leu Gly Arg Gly Val Phe Val Asn Ala Ser Gly Leu Arg ctg ctc gat cta tca tct aac acg ttg cgg gcg ctt ggc cgc cac gac Leu Leu Asp Leu Ser Ser Asn Thr Leu Arg Ala Leu Gly Arg His Asp 115 120 ctc gac ggg ctg ggg gcg ctg gag aag ctg ctt ctg ttc aat aac cgc 432 Leu Asp Gly Leu Gly Ala Leu Glu Lys Leu Leu Leu Phe Asn Asn Arg 130 135 ttg gtg cac ttg gac gag cat gcc ttc cac ggc ctg cgc gcg ctc agc Leu Val His Leu Asp Glu His Ala Phe His Gly Leu Arg Ala Leu Ser

145	15		155		160
cat ctc tac His Leu Tyr	c ctg ggc tgo c Leu Gly Cy: 165	c aac gaa s Asn Glu	ctc gcc tcg Leu Ala Ser 170	ttc tcc ttc Phe Ser Phe	gac cac 528 Asp His 175
ctg cac ggt Leu His Gly	ctg agc gcd Leu Ser Ala 180	acc cac Thr His	ctg ctt act Leu Leu Thr 185	ctg gac ctc Leu Asp Leu 190	tcc tcc 576 Ser Ser
aac cgg ctg Asn Arg Leu 195	GLY His Ile	tcc gta Ser Val 200	cct gag ctg Pro Glu Leu	gcc gcg ctg Ala Ala Leu 205	ccg gcc 624 Pro Ala
ttc ctc aag Phe Leu Lys 210	aac ggc cto Asn Gly Leu	tac ttg Tyr Leu 215	cac aac aac His Asn Asn	cct ttg cct Pro Leu Pro 220	tgc gac 672 Cys Asp
tgc cgc ctc Cys Arg Leu 225	tac cac cto Tyr His Leu 230	ı Leu Gln	cgc tgg cac Arg Trp His 235	cag cgg ggc Gln Arg Gly	ctg agc 720 Leu Ser 240
gcc gtg cgc Ala Val Arg	gac ttt gcg Asp Phe Ala 245	r cgc gag Arg Glu	tac gta tgc Tyr Val Cys 250	ttg gcc ttc Leu Ala Phe	aag gta 768 Lys Val 255
ccc gcg tcc Pro Ala Ser	cgc gtg cgc Arg Val Arg 260	Phe Phe	cag cac agc Gln His Ser 265	cgc gtc ttt Arg Val Phe 270	gag aac 816 Glu Asn
tgc tcg tcg Cys Ser Ser 275	gcc cca gct Ala Pro Ala	ctt ggc Leu Gly 280	cta gag cgg Leu Glu Arg	ccg gaa gag Pro Glu Glu 285	cac ctg 864 His Leu
tac gcg ctg Tyr Ala Leu 290	gtg ggt cgg Val Gly Arg	tcc ctg . Ser Leu . 295	agg ctt tac Arg Leu Tyr	tgc aac acc Cys Asn Thr 300	agc gtc 912 Ser Val
ccg gcc atg Pro Ala Met 305	cgc att gcc Arg Ile Ala 310	tgg gtt : Trp Val :	tcg ccg cag Ser Pro Gln 315	cag gag ctt Gln Glu Leu	ctc agg 960 Leu Arg 320
gcg cca gga Ala Pro Gly	tcc cgc gat Ser Arg Asp 325	ggc agc a	atc gcg gtg Ile Ala Val 330	ctg gcc gac Leu Ala Asp	ggc agc 1008 Gly Ser 335
ttg gcc ata Leu Ala Ile	ggc aac gta Gly Asn Val 340	Gin Glu G	cag cat gcg Gln His Ala 345	gga ctc ttc Gly Leu Phe 350	gtg tgc 1056 Val Cys
ctg gcc act Leu Ala Thr 355	ggg ccc cgc Gly Pro Arg	ctg cac c Leu His H 360	cac aac cag His Asn Gln	acg cac gag Thr His Glu ' 365	tac aac 1104 Tyr Asn
gtg agc gtg Val Ser Val 370	cac ttt ccg His Phe Pro	cgc cca g Arg Pro G 375	lu Pro Glu	gct ttc aac a Ala Phe Asn : 380	aca ggc 1152 Thr Gly
ttc acc aca Phe Thr Thr: 385	ctg ctg ggc Leu Leu Gly 390	tgt gcc g Cys Ala V	tg ggc ctt al Gly Leu 395	gtg ctc gtg o Val Leu Val I	ctg ctc 1200 Leu Leu 400

tac Tyr	ctg Leu	ttc Phe	gcc Ala	cca Pro 405	ccc Pro	tgc Cys	cgc Arg	tgc Cys	tgc Cys 410	cgc Arg	cgt Arg	gcc Ala	tgc Cys	cgc Arg 415	tgc Cys	1248
cgc Arg	cgc Arg	tgg Trp	ccc Pro 420	caa Gln	aca Thr	ccc Pro	agc Ser	ccg Pro 425	ctc Leu	caa Gln	gag Glu	ctg Leu	agc Ser 430	gca Ala	cag Gln	1296
tcc Ser	tca Ser	gta Val 435	ctc Leu	agc Ser	acc Thr	aca Thr	ccg Pro 440	cca Pro	gac Asp	gca Ala	ccc Pro	agc Ser 445	cgc Arg	aag Lys	gcc Ala	1344
agc Ser	gtc Val 450	cac His	aag Lys	cac His	gta Val	gtc Val 455	ttt Phe	ctg Leu	gag Glu	cca Pro	ggc Gly 460	cgg Arg	agg Arg	Gly ggc	ctc Leu	1392
aat Asn 465	gly	cgc Arg	gtg Val	cag Gln	ctg Leu 470	gca Ala	gta Val	gct Ala	gag Glu	gaa Glu 475	ttc Phe	gat Asp	ctc Leu	tac Tyr	aac Asn 480	1440
cct Pro	gga Gly	ggc Gly	ctg Leu	cag Gln 485	ctg Leu	aag Liys	gct Ala	ggc	tct Ser 490	gag Glu	tcc Ser	gcc Ala	agc Ser	tcc Ser 495	ata Ile	1488
	tcc Ser				_											1512
<210	ı															
	.> 50	4														
<212	> PR	2 <b>T</b>														

<213> Homo sapiens

<400> 6

Met Thr Trp Leu Val Leu Leu Gly Thr Leu Leu Cys Met Leu Arg Val

Gly Leu Gly Thr Pro Asp Ser Glu Gly Phe Pro Pro Arg Ala Leu His

Asn Cys Pro Tyr Lys Cys Ile Cys Ala Ala Asp Leu Leu Ser Cys Thr

Gly Leu Gly Leu Gln Asp Val Pro Ala Glu Leu Pro Ala Ala Thr Ala

Asp Leu Asp Leu Ser His Asn Ala Leu Gln Arg Leu Arg Pro Gly Trp

Leu Ala Pro Leu Phe Gln Leu Arg Ala Leu His Leu Asp His Asn Glu 85

Leu Asp Ala Leu Gly Arg Gly Val Phe Val Asn Ala Ser Gly Leu Arg 105

Leu Leu Asp Leu Ser Ser Asn Thr Leu Arg Ala Leu Gly Arg His Asp 115

Leu Asp Gly Leu Gly Ala Leu Glu Lys Leu Leu Leu Phe Asn Asn Arg

	130					135					140				
Leu 145	Val	His	Leu	Asp	Glu 150	His	Ala	Phe	His	Gly 155	Leu	Arg	Ala	Leu	Ser 160
His	Leu	Tyr	Leu	Gly 165	Cys	Asn	Glu	Leu	Ala 170	Ser	Phe	Ser	Phe	Asp 175	His
Leu	His	Gly	Leu 180		Ala	Thr	His	Leu 185	Leu	Thr	Leu	Asp	Leu 190	Ser	Ser
Asn	Arg	Leu 195	Gly	His	Ile	Ser	Val 200	Pro	Glu	Leu	Ala	Ala 205	Leu	Pro	Ala
Phe	Leu 210	Lys	Asn	Gly	Leu	Tyr 215	Leu	His	Asn	Asn	Pro 220	Leu	Pro	Cys	Asp
Cys 225	Arg	Leu	Tyr	His	Leu 230	Leu	Gln	Arg	Trp	His 235	Gln	Arg	Gly	Leu	Ser 240
Ala	Val	Arg	Asp	Phe 245	Ala	Arg	Glu	Tyr	Val 250	Cys	Leu	Ala	Phe	Lys 255	Val
Pro	Ala	Ser	Arg 260	Val	Arg	Phe	Phe	Gln 265	His	Ser	Arg	Val	Phe 270	Glu	Asn
Cys	Ser	Ser 275	Ala	Pro	Ala	Leu	Gly 280	Leu	Glu	Arg	Pro	Glu 285	Glu	His	Leu
Tyr	Ala 290	Leu	Val	Gly	Arg	Ser 295	Leu	Arg	Leu	Tyr	300 300	Asn	Thr	Ser	Val
Pro 305	Ala	Met	Arg	Ile	Ala 310	Trp	Val	Ser	Pro	Gln 315	Gln	Glu	Leu	Leu	Arg 320
Ala	Pro	Gly	Ser	Arg 325	Asp	Gly	Ser	Ile	Ala 330	Val	Leu	Ala	Asp	Gly 335	Ser
Leu	Ala	Ile	Gly 340	Asn	Val	Gln	Glu	Gln 345	His	Ala	Gly	Leu	Phe 350	Val	Cys
Leu	Ala	Th <i>r</i> 355	Gly	Pro	Arg	Leu	His 360	His	Asn	Gln	Thr	His 365	Glu	Tyr	Asn
Val	Ser 370	Val	His	Phe	Pro	Arg 375	Pro	Glu	Pro	Glu	Ala 380	Phe	Asn	Thr	Gly
Phe 385	Thr	Thr	Leu	Leu	Gly 390	Cys	Ala	Val	Gly	Leu 395	Val	Leu	Val	Leu	Leu 400
Tyr	Leu	Phe	Ala	Pro 405	Pro	Cys	Arg	Cys	Cys 410	Arg	Arg	Ala	Cys .	Arg 415	
Arg	Arg	Trp	Pro 420	Gln	Thr	Pro	Ser	Pro 425	Leu	Gln	Glu	Leu	Ser 430	Ala	Gln
Ser	Ser	Val 435	Leu	Ser	Thr	Thr	Pro 440	Pro	Asp	Ala	Pro	Ser 445	Arg	ГЛЗ	Ala
Ser	Val 450	His	Lys	His	Val	Val 455	Phe	Leu	Glu	Pro	Gly 460	Arg	Arg	Gly	Leu

Asn Gly Arg Val Gln Leu Ala Val Ala Glu Glu Phe Asp Leu Tyr Asn

Pro Gly Gly Leu Gln Leu Lys Ala Gly Ser Glu Ser Ala Ser Ser Ile Gly Ser Glu Gly Pro Met Thr Thr 500 <210> 7 <211> 1827 <212> DNA <213> Artificial Sequence <220> <223> Description of Artificial Sequence: Ig-fusion <220> <221> CDS <222> (1)..(1827) <400> 7 atg caa ccc cag cgt gac ctg cga ggc ctc tgg ctc ctg ctc tcc Met Gln Pro Gln Arg Asp Leu Arg Gly Leu Trp Leu Leu Leu Ser 96 gtg ttc ctg ctt ctc ttt gag gta gcc agg gcc ggc cga tct gtg gtt Val Phe Leu Leu Phe Glu Val Ala Arg Ala Gly Arg Ser Val Val 20 agt tgt ccc gcc aac tgc ctg tgc gcc agc aac atc ctc agc tgc tcc 144 Ser Cys Pro Ala Asn Cys Leu Cys Ala Ser Asn Ile Leu Ser Cys Ser 35 aag cag cag ctg ccc aat gtg ccc caa tct ttg ccc agc tac aca gca 192 Lys Gln Gln Leu Pro Asn Val Pro Gln Ser Leu Pro Ser Tyr Thr Ala 50 ctg ctg gac ctc agc cac aac aac ttg agc agg ctg cgg gcc gag tgg Leu Leu Asp Leu Ser His Asn Asn Leu Ser Arg Leu Arg Ala Glu Trp 65 70 acc ecc acc egg etg acc aac etg cac tee etg etg age cac aac Thr Pro Thr Arq Leu Thr Asn Leu His Ser Leu Leu Leu Ser His Asn 85 cac ctq aac ttc atc tcc tcc gag gcc ttc gtc ccc gta ccc aac ctt 336 His Leu Asn Phe Ile Ser Ser Glu Ala Phe Val Pro Val Pro Asn Leu 100 agg tac ttg gac ctc tcc tcc aac cat ctt cac acg ctg gat gag ttc 384 Arg Tyr Leu Asp Leu Ser Ser Asn His Leu His Thr Leu Asp Glu Phe 120 115 ctg ttc agc gac ctg cag gcg ctg gaa gtg ctg ttg ctc tac aat aac Leu Phe Ser Asp Leu Gln Ala Leu Glu Val Leu Leu Tyr Asn Asn 130

cac His	: ITE	gto Val	g gto Val	gtg Val	gac Asp 150	Arg	aat Asn	gcc	ttt Phe	gag Glu 155	1 Asp	atg Met	gco Ala	cag Gln	ctg Leu 160	480
caç Glr	aaa Lys	cto Lev	tac Tyr	tta Leu 165	Ser	cag Gln	aat Asn	Cag Gln	ato Ile 170	Ser	cgc Arg	ttt Phe	cct Pro	gtg Val 175	gaa Glu	528
ct <u>e</u> Leu	ato Ile	aag Lys	gat Asp 180	Gly	aac Asn	aaa Lys	tta Leu	Pro 185	Lys	ctg Leu	atg Met	ctc Leu	ttg Leu 190	Asp	ctg Leu	576
tcc Ser	tcc Ser	aac Asn 195	Lys	ctg Lev	aag Lys	aag Lys	ttg Leu 200	Pro	ctg Leu	act Thr	gac Asp	ctg Leu 205	cag Gln	aaa Lys	ttg Leu	624
cca Pro	gcc Ala 210	Trp	gto Val	aag Lys	aat Asn	999 Gly 215	cta Leu	tac Tyr	ctg Leu	cat His	aac Asn 220	aac Asn	ccc Pro	ttg Leu	gag Glu	672
tgc Cys 225	Asp	tgc Cys	aag Lys	ctc Leu	tac Tyr 230	cag Gln	ctc Leu	ttt Phe	tcg Ser	cac His 235	tgg Trp	cag Gln	tac Tyr	cgg Arg	cag Gln 240	720
ctg Leu	agc Ser	tct Ser	gtg Val	atg Met 245	gac Asp	ttc Phe	cag Gln	gag Glu	gac Asp 250	ctg Leu	tac Tyr	tgc Cys	atg Met	cac His 255	tcc Ser	768
aag Lys	aag Lys	ctg Leu	cac His 260	Asn	atc Ile	ttc Phe	agc Ser	ctg Leu 265	gat Asp	ttc Phe	ttc Phe	aat Asn	tgc Cys 270	agc Ser	gag Glu	816
tac Tyr	aag Lys	gaa Glu 275	agt Ser	gcc Ala	tgg Trp	gag Glu	gct Ala 280	cac His	ctg Leu	gga Gly	gac Asp	acc Thr 285	ttg Leu	acc Thr	atc Ile	864
agg Arg	tgt Cys 290	gac Asp	acc Thr	aaa Lys	cag Gln	caa Gln 295	ggc Gly	atg Met	acc Thr	aaa Lys	gtg Val 300	tgg Trp	gtg Val	tcc Ser	cca Pro	912
agc Ser 305	aat Asn	gaa Glu	cag Gln	gtg Val	cta Leu 310	agt Ser	cag Gln	gjå aaa	tcc Ser	aat Asn 315	ggc Gly	tcg Ser	gtg Val	agc Ser	gtg Val 320	960
agg Arg	aat Asn	ggc Gly	gac Asp	ctt Leu 325	ttt Phe	ttt Phe	aaa Lys	aag Lys	gtg Val 330	cag Gln	gtc Val	gag Glu	gat Asp	999 935 335	ggt Gly	1008
gtg Val	tat Tyr	acc Thr	tgt Cys 340	tac Tyr	gcc Ala	atg Met	gly aaa	gag Glu 345	act Thr	ttc Phe	aac Asn	gag Glu	aca Thr 350	ctg Leu	tct Ser	1056
gtg Val	gag Glu	ttg Leu 355	aaa Lys	gtg Val	tat Tyr	aac Asn	ttc Phe 360	acc Thr	ttg Leu	cac His	gga Gly	cac His 365	cat His	gac Asp	acc Thr	1104
ctc Leu	aac Asn 370	gga Gly	tcc Ser	gag Glu	Val	ctg Leu 375	ttc Phe	cag Gln	ggc Gly	ccc Pro	aaa Lys 380	tct Ser	tgt Cys	gac <sup>.</sup> Asp	aaa Lys	1152

	cac His											1200
	gtc Val						_		_			1248
	acc Thr		_	_		 	_	 _		-		1296
	gag Glu											1344
	aag Lys 450											1392
	agc Ser											1440
	aag Lys											1488
	atc Ile		_		_	_	_	_			acc Thr	1536
	ccc Pro											1584
	ctg Leu 530											1632
	aat Asn											1680
	tcc Ser											1728
	agg Arg											1776
	ctg Leu											1824
aaa Lys												1827

<210> 8

<211> 609

<212> PRT

<213> Artificial Sequence

<223> Description of Artificial Sequence: Ig-fusion protein

<400> 8

Met Gln Pro Gln Arg Asp Leu Arg Gly Leu Trp Leu Leu Leu Ser 1 5 10 15

Val Phe Leu Leu Phe Glu Val Ala Arg Ala Gly Arg Ser Val Val 20 25 30

Ser Cys Pro Ala Asn Cys Leu Cys Ala Ser Asn Ile Leu Ser Cys Ser 35 40 45

Lys Gln Gln Leu Pro Asn Val Pro Gln Ser Leu Pro Ser Tyr Thr Ala 50 55 60

Leu Leu Asp Leu Ser His Asn Asn Leu Ser Arg Leu Arg Ala Glu Trp 65 70 75 80

Thr Pro Thr Arg Leu Thr Asn Leu His Ser Leu Leu Leu Ser His Asn
85 90 95

His Leu Asn Phe Ile Ser Ser Glu Ala Phe Val Pro Val Pro Asn Leu 100 105 110

Arg Tyr Leu Asp Leu Ser Ser Asn His Leu His Thr Leu Asp Glu Phe 115 120 125

Leu Phe Ser Asp Leu Gln Ala Leu Glu Val Leu Leu Leu Tyr Asn Asn 130 135 140

His Ile Val Val Val Asp Arg Asn Ala Phe Glu Asp Met Ala Gln Leu 145 150 155 160

Gln Lys Leu Tyr Leu Ser Gln Asn Gln Ile Ser Arg Phe Pro Val Glu 165 170 175

Leu Ile Lys Asp Gly Asn Lys Leu Pro Lys Leu Met Leu Leu Asp Leu 180 185 190

Ser Ser Asn Lys Leu Lys Lys Leu Pro Leu Thr Asp Leu Gln Lys Leu 195 200 205

Pro Ala Trp Val Lys Asn Gly Leu Tyr Leu His Asn Asn Pro Leu Glu 210 215 220

Cys Asp Cys Lys Leu Tyr Gln Leu Phe Ser His Trp Gln Tyr Arg Gln 225 230 235 240

Leu Ser Ser Val Met Asp Phe Gln Glu Asp Leu Tyr Cys Met His Ser 245 250 255

Lys Lys Leu His Asn Ile Phe Ser Leu Asp Phe Phe Asn Cys Ser Glu 260 265 270

Tyr Lys Glu Ser Ala Trp Glu Ala His Leu Gly Asp Thr Leu Thr Ile 275 280 285

Arg Cys Asp Thr Lys Gln Gln Gly Met Thr Lys Val Trp Val Ser Pro 290 295 300

Ser Asn Glu Gln Val Leu Ser Gln Gly Ser Asn Gly Ser Val Ser Val 305 310 315 320

Arg Asn Gly Asp Leu Phe Phe Lys Lys Val Gln Val Glu Asp Gly Gly 325 330 335

Val Tyr Thr Cys Tyr Ala Met Gly Glu Thr Phe Asn Glu Thr Leu Ser 340 345 350

Val Glu Leu Lys Val Tyr Asn Phe Thr Leu His Gly His His Asp Thr 355 360 365

Leu Asn Gly Ser Glu Val Leu Phe Gln Gly Pro Lys Ser Cys Asp Lys 370 380

Thr His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu Gly Gly Pro 385 390 395 400

Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr Leu Met Ile Ser 405 · 410 415

Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser His Glu Asp 420 430

Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val Glu Val His Asn 435 440 445

Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser Thr Tyr Arg Val 450 455 460

Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu Asn Gly Lys Glu 465 470 475 480

Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala Pro Ile Glu Lys
485 490 495

Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro Gln Val Tyr Thr 500 505 510

Leu Pro Pro Ser Arg Asp Glu Leu Thr Lys Asn Gln Val Ser Leu Thr 515 520 525

Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu 530 540

Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu 545 550 555 560

Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys
565 570 575

Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu 580 585 590

Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly
595 600 605

Lys

```
<210> 9
<211> 1920
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Ig-fusion
      protein
<220>
<221> CDS
<222> (1)..(1920)
<400> 9
atg tog tta agg tto cac aca ctg ccc acc ctg cct aga gct gtc aaa
                                                                   48
Met Ser Leu Arg Phe His Thr Leu Pro Thr Leu Pro Arg Ala Val Lys
ccg ggt tgc aga gag ctg ctg tgt ctg ttg gtg atc gca gtg atg gtg
                                                                   96
Pro Gly Cys Arg Glu Leu Leu Cys Leu Leu Val Ile Ala Val Met Val
age eee age gee tea gga atg tge eee act get tge ate tgt gee ace
                                                                   144
Ser Pro Ser Ala Ser Gly Met Cys Pro Thr Ala Cys Ile Cys Ala Thr
         35
                              40
                                                                   192
gac att gtc agc tgc acc aac aaa aac cta tct aag gtg ccc ggg aac
Asp Ile Val Ser Cys Thr Asn Lys Asn Leu Ser Lys Val Pro Gly Asn
                                              60
                          55
     50
ctt ttc aga ctg att aaa aga ctg gat ctg agc tat aac aga atc gga
                                                                   240
Leu Phe Arg Leu Ile Lys Arg Leu Asp Leu Ser Tyr Asn Arg Ile Gly
                      70
 65
ctg ttg gat gcc gac tgg atc ccg gtg tcg ttt gtc aag ctg agc acc
                                                                   288
Leu Leu Asp Ala Asp Trp Ile Pro Val Ser Phe Val Lys Leu Ser Thr
                                      90
                  85
tta att ctt cgc cac aac atc acc agc atc tcc acg ggc agt ttc
                                                                   336
Leu Ile Leu Arg His Asn Asn Ile Thr Ser Ile Ser Thr Gly Ser Phe
                                 105
             100
tec aca acc cca aat tta aag tgt ctg gac tta tca tcc aat agg ctg
                                                                    384
Ser Thr Thr Pro Asn Leu Lys Cys Leu Asp Leu Ser Ser Asn Arg Leu
                                                 125
         115
aag tog gta aag agt goo aca tto caa gag otg aag got otg gaa gta
Lys Ser Val Lys Ser Ala Thr Phe Gln Glu Leu Lys Ala Leu Glu Val
                         135
     130
ctg ctg ctg tac aac aac cac att tcc tat ctg gac ccc gca gcg ttc
Leu Leu Leu Tyr Asn Asn His Ile Ser Tyr Leu Asp Pro Ala Ala Phe
                                                              160
                                         155
                     150
145
 ggg ggg ctt tcc cac ttg cag aaa ctc tat ctg agt ggg aac ttt ctc
 Gly Gly Leu Ser His Leu Gln Lys Leu Tyr Leu Ser Gly Asn Phe Leu
                 165
                                     170
```

aca Thr	cag Gln	tto Phe	cct Pro	) Met	gat : Asp	ttg Leu	tat Tyr	act Thr 185	Gly	agg Arg	tto Phe	aag Lys	cto Leu 190	ı Ala	gat Asp	576
ctg Leu	aca Thr	ttt Phe 195	Lev	gat Asp	gtt Val	tcc Ser	tat Tyr 200	Asn	cga Arg	ato Ile	cct Pro	Ser 205	: Ile	ccg Pro	atg Met	624
cac His	cat His 210	Ile	aac Asn	tta Leu	gtg Val	ccg Pro 215	Gly	aga Arg	cag Gln	ctg Leu	aga Arg 220	Gly	ato Ile	tac Tyr	ctt Leu	672
cac His 225	Gly	aac Asn	cca Pro	ttt Phe	gta Val 230	Cys	gac Asp	tgt Cys	tct Ser	ctg Leu 235	Tyr	tcg Ser	ttg Leu	ctg Leu	atc Ile 240	720
ttt Phe	tgg Trp	tac Tyr	cgt Arg	agg Arg 245	His	ttt Phe	agc Ser	tcc Ser	gtg Val 250	atg Met	gat Asp	ttt Phe	aag Lys	aat Asn 255	gac Asp	768
tat Tyr	acc Thr	tgt Cys	cgc Arg 260	ctg Leu	tgg Trp	tct Ser	gac Asp	tcc Ser 265	agg Arg	cac His	tcc Ser	cac His	cag Gln 270	ctg Leu	cag Gln	816
ctg Leu	ctc Leu	cag Gln 275	gag Glu	agc Ser	ttt Phe	ctg Leu	aac Asn 280	tgt Cys	tct Ser	tac Tyr	agc Ser	gtt Val 285	atc Ile	aac Asn	ggc Gly	864
tcc Ser	ttc Phe 290	cac His	gca Ala	ctt Leu	Gly	ttt Phe 295	atc Ile	cac His	gag Glu	gct Ala	cag Gln 300	gtt Val	gly aaa	gag Glu	agg Arg	912
gcg Ala 305	atc Ile	gtc Val	cac His	tgt Cys	gac Asp 310	agc Ser	aag Lys	act Thr	ggc	aat Asn 315	gga Gly	aat Asn	act	gat Asp	ttc Phe 320	960
atc Ile	tgg Trp	gtc Val	ggt Gly	ccc Pro 325	gat Asp	aac Asn	agg Arg	ctg Leu	ctg Leu 330	gag Glu	cca Pro	gat Asp	aaa Lys	gac Asp 335	atg Met	1008
ej aaa	aac Asn	ttt Phe	cgt Arg 340	gtg Val	ttt Phe	tac Tyr	aac Asn	gga Gly 345	agt Ser	ctg Leu	gtc Val	ata Ile	gag Glu 350	aac Asn	cct Pro	1056
ggc	ttt Phe	gag Glu 355	gac Asp	gcc Ala	gjå aaa	gta Val	tat Tyr 360	tct Ser	tgt Cys	atc Ile	gca Ala	atg Met 365	aac Asn	agg Arg	cag Gln	1104
cgg Arg	ctg Leu 370	tta Leu	aac Asn	gag Glu	acg Thr	gtg Val 375	gat Asp	atc Ile	atg Met	atc Ile	aac Asn 380	gtg Val	agc Ser	aat Asn	ttc Phe	1152
acc Thr 385	ata Ile	aac Asn	aga Arg	tcc Ser	cac His 390	gcc Ala	cac His	gag Glu	gcg Ala	gcg Ala 395	gcc Ala	gcg Ala	gat Asp	ccc Pro	atc Ile 400	1200
gaa Glu	ggt Gly	cgt Arg	ggt Gly	ggt Gly 405	ggt Gly	ggt Gly	ggt Gly	Asp	ccc Pro 410	aaa Lys	tct Ser	tgt Cys	Asp	aaa Lys 415	cct Pro	1248

cac	aca Thr	tgc Cys	cca Pro 420	Pro	tgc Cys	cca Pro	gca Ala	cct Pro 425	Glu	cto Leu	cto Leu	Gly 999	gga Gly 430	Pro	g tca Ser	1296
gto Val	tto Phe	Leu 435	Phe	Pro	cca Pro	aaa Lys	Pro 440	Lys	gac Asp	acc Thr	ctc Leu	atg Met 445	Ile	tco Ser	cgg Arg	1344
acc Thr	Pro 450	Glu	gtc Val	aca Thr	tgc Cys	gtg Val 455	gtg Val	gtg Val	gac Asp	gtg Val	ago Ser 460	His	gaa Glu	gac Asp	cct Pro	1392
gag Glu 465	val	aag Lys	ttc Phe	aac Asn	tgg Trp 470	tac Tyr	gtg Val	gac	ggc	gtg Val 475	gag Glu	gtg Val	cat His	aat Asn	gcc Ala 480	1440
гÀЗ	Thr	гÀз	ccg Pro	Arg 485	Glu	Glu	Gln	Tyr	Asn 490	Ser	Thr	Tyr	Arg	Val 495	Val	1488
ser	vaı	Leu	acc Thr 500	Val	Leu	His	Gln	Asp 505	Trp	Leu	Asn	Gly	Lys 510	Glu	Tyr	1536
гуз	Cys	Ьуз 515	gtc Val	Ser	Asn	Lys	Ala 520	Leu	Pro	Ala	Pro	Ile 525	Glu	Lys	Thr	1584
TTE	530	Lys	gcc Ala	Lys	Gly	Gln 535	Pro	Arg	Glu	Pro	Gln 540	Val	Tyr	Thr	Leu	1632
Pro 545	Pro	Ser	cgg Arg	Asp	Glu 550	Leu	Thr	Lys	Asn	Gln 555	Val	Ser	Leu	Thr	Cys 560	1680
Leu	vai	тÃв	Gly	Phe 565	Tyr	Pro	Ser	Asp	Ile 570	Ala	Val	Glu	Trp	Glu 575	Ser	1728
Asn	GIÀ	GIn	ccg Pro 580	Glu	Asn	Asn	Tyr	Lys 585	Thr	Thr	Pro	Pro	Val 590	Leu	Asp	1776
ser	Asp	595	tcc Ser	Phe	Phe	Leu	Tyr 600	Ser	ГÀЗ	Leu	Thr	Val 605	Asp	Lys	Ser	1824
Arg	610	GIN	cag Gln	GIA	Asn	Val 615	Phe	Ser	Cys	Ser	Val 620	Met	His	Glu	Ala	1872
ctg Leu 625	cac His	aac Asn	cac His	Tyr	acg Thr 630	cag Gln	aag Lys	agc Ser	Leu	tcc Ser 635	ctg Leu	tct Ser	ccg Pro	ggt Gly	aaa Lys 640	1920

<sup>&</sup>lt;210> 10 <211> 640 <212> PRT <213> Artificial Sequence

<223> Description of Artificial Sequence: Ig-fusion
 protein

<400> 10 Met Ser Leu Arg Phe His Thr Leu Pro Thr Leu Pro Arg Ala Val Lys Pro Gly Cys Arg Glu Leu Leu Cys Leu Leu Val Ile Ala Val Met Val Ser Pro Ser Ala Ser Gly Met Cys Pro Thr Ala Cys Ile Cys Ala Thr Asp Ile Val Ser Cys Thr Asn Lys Asn Leu Ser Lys Val Pro Gly Asn Leu Phe Arg Leu Ile Lys Arg Leu Asp Leu Ser Tyr Asn Arg Ile Gly Leu Leu Asp Ala Asp Trp Ile Pro Val Ser Phe Val Lys Leu Ser Thr Leu Ile Leu Arg His Asn Asn Ile Thr Ser Ile Ser Thr Gly Ser Phe Ser Thr Thr Pro Asn Leu Lys Cys Leu Asp Leu Ser Ser Asn Arg Leu Lys Ser Val Lys Ser Ala Thr Phe Gln Glu Leu Lys Ala Leu Glu Val 130 Leu Leu Tyr Asn Asn His Ile Ser Tyr Leu Asp Pro Ala Ala Phe 150 155 Gly Gly Leu Ser His Leu Gln Lys Leu Tyr Leu Ser Gly Asn Phe Leu 165 170 Thr Gln Phe Pro Met Asp Leu Tyr Thr Gly Arg Phe Lys Leu Ala Asp Leu Thr Phe Leu Asp Val Ser Tyr Asn Arg Ile Pro Ser Ile Pro Met His His Ile Asn Leu Val Pro Gly Arg Gln Leu Arg Gly Ile Tyr Leu His Gly Asn Pro Phe Val Cys Asp Cys Ser Leu Tyr Ser Leu Leu Ile Phe Trp Tyr Arg Arg His Phe Ser Ser Val Met Asp Phe Lys Asn Asp Tyr Thr Cys Arg Leu Trp Ser Asp Ser Arg His Ser His Gln Leu Gln Leu Leu Gln Glu Ser Phe Leu Asn Cys Ser Tyr Ser Val Ile Asn Gly Ser Phe His Ala Leu Gly Phe Ile His Glu Ala Gln Val Gly Glu Arg 300

Ala 305	Ile	Val	His	Сув	Asp 310	Ser	Lys	Thr	Gly	Asn 315	Gly	Asn	Thr	Asp	Phe 320
Ile	Trp	Val	Gly	Pro 325	Asp	Asn	Arg	Leu	Leu 330	Glu	Pro	Asp	Lys	Asp 335	Met
Gly	Asn	Phe	Arg 340	Val	Phe	Tyr	Asn	Gly 345	Ser	Leu	Val	Ile	Glu 350	Asn	Pro
Gly	Phe	Glu 355	Asp	Ala	Gly	Val	Tyr 360	Ser	Cys	Ile	Ala	Met 365	Asn	Arg	Gln
Arg	Leu 370	Leu	Asn	Glu	Thr	Val 375	Asp	Ile	Met	Ile	Asn 380	Val	Ser	Asn	Phe
Thr 385	Ile	Asn	Arg	Ser	His 390	Ala	His	Glu	Ala	Ala 395	Ala	Ala	Asp	Pro	Ile 400
Glu	Gly	Arg	Gly	Gly 405	Gly	Gly	Gly	Asp	Pro 410	Lys	Ser	Сув	Asp	Lys 415	Pro
His	Thr	Cys	Pro 420	Pro	Cys	Pro	Ala	Pro 425	Glu	Leu	Leu	Gly	Gly 430	Pro	Ser
Val	Phe	Leu 435	Phe	Pro	Pro	Lys	Pro 440	ГЛЗ	Asp	Thr	Leu	Met 445	Ile	Ser	Arg
Thr	Pro 450	Glu	Val	Thr	Cys	Val 455	Val	Val	Asp	Val	Ser 460	His	Glu	Asp	Pro
Glu 465	Val	Lys	Phe	Asn	Trp 470	Tyr	Val	Asp	Gly	Val 475	Glu	Val	His	Asn	Ala 480
ГÀЗ	Thr	Lys	Pro	Arg 485	Glu	Glu	Gln	Tyr	Asn 490	Ser	Thr	Tyr	Arg	Val 495	Val
Ser	Val	Leu	Thr 500	Val	Leu	His	Gln	Asp 505	Trp	Leu	Asn	Gly	Lys 510	Glu	Tyr
Lys	Сув	Lys 515	Val	Ser	Asn	Lys	Ala 520	Leu	Pro	Ala	Pro	Ile 525	Glu	Lys	Thr
Ile	Ser 530	Lys	Ala	Lys	Gly	Gln 535	Pro	Arg	Glu	Pro	Gln 540	Val	Tyr	Thr	Leu
Pro 545	Pro	Ser	Arg	Asp	Glu 550	Leu	Thr	Lys	Asn	Gln 555	Val	Ser	Leu	Thr	Cys 560
Leu	Val	Lys	Gly	Phe 565	Tyr	Pro	Ser	Asp	Ile 570	Ala	Val	Glu	Trp	Glu 575	Ser
Asn	Gly	Gln	Pro 580	Glu	Asn	Asn	Tyr	Lys 585	Thr	Thr	Pro	Pro	Val 590	Leu	qeA
Ser	Asp	Gly 595	Ser	Phe	Phe	Leu	Tyr 600	Ser	Lys	Leu	Thr	Val 605	Asp	Lys	Ser
Arg	Trp 610	Gln	Gln	Gly	Asn	Val 615	Phe	Ser	Cys	Ser	Val 620	Met	His	Glu	Ala

Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser Leu Ser Pro Gly Lys 625 630 635 640

<210> 11

<211> 1887 <212> DNA <213> Artificial Sequence <223> Description of Artificial Sequence: Ig-fusion protein <220> <221> CDS <222> (1)..(1887) <400> 11 48 atg gcc tgg cta gtg cta tca ggt ata cta cta tgc atg ttg ggt gct Met Ala Trp Leu Val Leu Ser Gly Ile Leu Leu Cys Met Leu Gly Ala gga ttg ggc act tca gac ttg gag gat gtt ctg cct cct gct ccc cac 96 Gly Leu Gly Thr Ser Asp Leu Glu Asp Val Leu Pro Pro Ala Pro His aac tgc ccc gat ata tgc atc tgt gct gcc gat gtg ttg agc tgt gcg i 144 Asn Cys Pro Asp Ile Cys Ile Cys Ala Ala Asp Val Leu Ser Cys Ala ggc cgt ggg tta cag gac ttg ccg gta gca ctg cct acc act gct gca 192 Gly Arg Gly Leu Gln Asp Leu Pro Val Ala Leu Pro Thr Thr Ala Ala gaa ctc gat ttg agc cac aac gca ctc aaa cgc ctg cac ccg ggg tgg 240 Glu Leu Asp Leu Ser His Asn Ala Leu Lys Arg Leu His Pro Gly Trp tta geg eec etc tee egg etg egt gee ttg eac eta gge tat aat aag 288 Leu Ala Pro Leu Ser Arg Leu Arg Ala Leu His Leu Gly Tyr Asn Lys ctq qaa gtc ctg ggc cat ggt gcg ttc acc aat gcc agt ggc ctg agg 336 Leu Glu Val Leu Gly His Gly Ala Phe Thr Asn Ala Ser Gly Leu Arg 105 aca ctt gac ctg tcc tct aat atg tta agg atg ctc cat acc cat gac 384 Thr Leu Asp Leu Ser Ser Asn Met Leu Arg Met Leu His Thr His Asp 115 120 ctg gat ggc ctg gag gag ctg gag aag tta ctt ctg ttc aat aac agc 432 Leu Asp Gly Leu Glu Glu Leu Glu Lys Leu Leu Phe Asn Asn Ser 135 130 ctg atg cac ttg gac ctg gat gcc ttc cag ggc ctg cgc atg ctt agc 480 Leu Met His Leu Asp Leu Asp Ala Phe Gln Gly Leu Arg Met Leu Ser 150 155 145 cac ctc tat ctc agc tgc aac gag ctc tcc tct ttc tct ttc aac cac 528 His Leu Tyr Leu Ser Cys Asn Glu Leu Ser Ser Phe Ser Phe Asn His

•																
				165					170					175		
Lei	g cac 1 His	ggt Gly	ctg Leu 180	Gly	, tta , Leu	acc Thr	e cgo	ctg Leu 185	ı Arg	g act g Thr	cto Lei	g gad 1 Asp	t cto Lev 190	ı Sei	tcc Ser	576
aac	tgg	, ctg	, aaa	cat	ato	tco	ato	cct	gag	, ttg	g gct	gca	a cte	g cca	a act	624
Asr	Trp	195		His	: Ile	: Ser	11e 200		Glu	Leu	a Ala	Ala 205		Pro	Thr	
tat Tyr	Leu 210	Lys	aac Asn	agg Arg	ctc Leu	tac Tyr 215	Leu	cac His	aac Asn	aac Asn	Pro 220	Let	g ccc i Pro	tgt Cys	gac Asp	672
tgo Cys 225	Ser	cto Leu	tac Tyr	cac His	Leu 230	Leu	cgg Arg	cgc Arg	tgg Trp	Cac His 235	Gln	g cgg	Gly Gly	ter	g agt Ser 240	720
gcc	ctg Leu	cat His	gat Asp	ttt Phe 245	Glu	cgc Arg	gag Glu	tac Tyr	aca Thr 250	Cys	ttg Leu	gto Val	ttt. Phe	aag Lys 255	gtg Val	768
tca Ser	gag Glu	tcc Ser	cga Arg 260	gtg Val	cgc Arg	ttt Phe	ttt Phe	gag Glu 265	cac His	agc Ser	cgg Arg	gtc Val	Phe 270	Lys	aac Asn	816
tgc Cys	tct Ser	gtg Val 275	Ala	gca Ala	gct Ala	cca Pro	ggc Gly 280	tta Leu	gag Glu	ctg Leu	cct Pro	gaa Glu 285	Glu	cag Gln	ctg Leu	864
cac His	gcg Ala 290	cag Gln	gtg Val	ggc	cag Gln	tcc Ser 295	ctg Leu	agg Arg	ctc Leu	ttc Phe	tgc Cys 300	aac Asn	acc Thr	agt Ser	gtg Val	912
cct Pro 305	gcc Ala	act Thr	cgg Arg	gtg Val	gcc Ala 310	tgg Trp	gtc Val	tcc Ser	ccg Pro	aag Lys 315	aat Asn	gag Glu	ctg Leu	ctt Leu	gtg Val 320	960
gcg Ala	cca Pro	gcc Ala	tct Ser	cag Gln 325	gat Asp	ggt Gly	agc Ser	atc Ile	gct Ala 330	gtg Val	ttg Leu	gct Ala	gat Asp	ggc Gly 335	agc Ser	1008
tta Leu	gcc Ala	ata Ile	ggc Gly 340	agg Arg	gtg Val	caa Gln	gag Glu	cag Gln 345	cac His	gca Ala	ggc Gly	gtc Val	ttt Phe 350	gtg Val	tgc Cys	1056
ctg Leu	gcc Ala	agt Ser 355	gly aaa	ccc Pro	cgc Arg	ctg Leu	cac His 360	cac His	aac Asn	cag Gln	aca Thr	ctt Leu 365	gag Glu	tac Tyr	aat Asn	1104
gtg Val	agt Ser 370	gtg Val	caa Gln	aag Lys	gct Ala	cgc Arg 375	ccc Pro	gag Glu	cca Pro	gag Glu	act Thr 380	ttc Phe	aac Asn	aca Thr	gcg Ala	1152
gcc Ala 385	gcg Ala	gat Asp	ccc Pro	atc Ile	gaa Glu 390	ggt Gly	cgt Arg	ggt Gly	ggt Gly	ggt Gly 395	ggt Gly	ggt Gly	gat Asp	Pro	aaa Lys 400	1200
tct Ser	tgt Cys	gac Asp	aaa Lys	cct Pro	cac His	aca Thr	tgc Cys	cca Pro	ccg Pro	tgc Cys	cca Pro	gca Ala	cct Pro	gaa Glu	ctc Leu	1248

	•			405	i				410					415		
ctç Lev	Gly 999	gga Gly	ccg Pro 420	Ser	gtc Val	ttc Phe	ctc Leu	tto Phe 425	Pro	cca Pro	aaa Lys	ccc	aag Lys 430	Asp	acc Thr	1296
cto Leu	atg Met	ato Ile 435	Ser	cgg	acc Thr	cct Pro	gag Glu 440	gtc Val	aca Thr	tgc Cys	gtg Val	gtg Val 445	gtg Val	gac Asp	gtg Val	1344
agc Ser	cac His 450	GIU	gac Asp	cct Pro	gag Glu	gtc Val 455	aag Lys	ttc Phe	aac Asn	tgg Trp	tac Tyr 460	gtg Val	gac Asp	ggc	gtg Val	1392
gag Glu 465	. vaı	cat His	aat Asn	gcc Ala	aag Lys 470	aca Thr	aag Lys	ccg Pro	cgg Arg	gag Glu 475	gag Glu	cag Gln	tac Tyr	aac Asn	agc Ser 480	1440
acg Thr	tac Tyr	cgg Arg	gtg Val	gtc Val 485	agc Ser	gtc Val	ctc Leu	Thr	gtc Val 490	ctg Leu	cac His	cag Gln	gac Asp	tgg Trp 495	ctg Leu	1488
aat Asn	ggc	aag Lys	gag Glu 500	tac Tyr	aag Lys	tgc Cys	aag Lys	gtc Val 505	tcc Ser	aac Asn	aaa Lys	gcc Ala	ctc Leu 510	cca Pro	gcc Ala	1536
ccc Pro	atc Ile	gag Glu 515	aaa Lys	acc Thr	atc Ile	tcc Ser	aaa Lys 520	gcc Ala	aaa Lys	gjå aaa	cag Gln	ccc Pro 525	cga Arg	gaa Glu	cca Pro	1584
cag Gln	gtg Val 530	tac Tyr	acc Thr	ctg Leu	ccc Pro	cca Pro 535	tcc Ser	cgg Arg	gat Asp	gag Glu	ctg Leu 540	acc Thr	aag Lys	aac Asn	cag Gln	1632
gtc Val 545	agc Ser	ctg Leu	acc Thr	tgc Cys	ctg Leu 550	gtc Val	aaa Lys	ggc Gly	ttc Phe	tat Tyr 555	ccc Pro	agc Ser	gac Asp	atc Ile	gcc Ala 560	1680
gtg Val	gag Glu	tgg Trp	gag Glu	agc Ser 565	aat Asn	GJÀ aaa	cag Gln	ccg Pro	gag Glu 570	aac Asn	aac Asn	tac Tyr	aag Lys	acc Thr 575	acg Thr	1728
cct Pro	ccc Pro	gtg Val	ctg Leu 580	gac Asp	tcc Ser	gac Asp	ggc Gly	tcc Ser 585	ttc Phe	ttc Phe	ctc Leu	tac Tyr	agc Ser 590	aag Lys	ctc Leu	1776
acc Thr	gtg Val	gac Asp 595	aag Lys	agc Ser	agg Arg	tgg Trp	cag Gln 600	cag Gln	gly aaa	aac Asn	gtc Val	ttc Phe 605	tca Ser	tgc Cys	tcc Ser	1824
gtg Val	atg Met 610	cat His	gag Glu	gct Ala	ctg Leu	cac His 615	aac Asn	cac His	tac Tyr	acg Thr	cag Gln 620	aag Lys	agc Ser	ctc Leu	tcc Ser	1872
			ggt Gly												-	1887

- <211> 629
- <212> PRT
- <213> Artificial Sequence
- <223> Description of Artificial Sequence: Ig-fusion protein

<400> 12

- Met Ala Trp Leu Val Leu Ser Gly Ile Leu Leu Cys Met Leu Gly Ala 1 5 10 15
- Gly Leu Gly Thr Ser Asp Leu Glu Asp Val Leu Pro Pro Ala Pro His
  20 25 30
- Asn Cys Pro Asp Ile Cys Ile Cys Ala Ala Asp Val Leu Ser Cys Ala 35 40 45
- Gly Arg Gly Leu Gln Asp Leu Pro Val Ala Leu Pro Thr Thr Ala Ala 50 55 60
- Glu Leu Asp Leu Ser His Asn Ala Leu Lys Arg Leu His Pro Gly Trp
  65 70 75 80
- Leu Ala Pro Leu Ser Arg Leu Arg Ala Leu His Leu Gly Tyr Asn Lys 85 90 95
- Leu Glu Val Leu Gly His Gly Ala Phe Thr Asn Ala Ser Gly Leu Arg
- Thr Leu Asp Leu Ser Ser Asn Met Leu Arg Met Leu His Thr His Asp 115 120 125
- Leu Asp Gly Leu Glu Glu Leu Glu Lys Leu Leu Leu Phe Asn Asn Ser 130 135 140
- Leu Met His Leu Asp Leu Asp Ala Phe Gln Gly Leu Arg Met Leu Ser 145 150 155 160
- His Leu Tyr Leu Ser Cys Asn Glu Leu Ser Ser Phe Ser Phe Asn His
  165 170 175
- Leu His Gly Leu Gly Leu Thr Arg Leu Arg Thr Leu Asp Leu Ser Ser 180 185 190
- Asn Trp Leu Lys His Ile Ser Ile Pro Glu Leu Ala Ala Leu Pro Thr 195 200 205
- Tyr Leu Lys Asn Arg Leu Tyr Leu His Asn Asn Pro Leu Pro Cys Asp 210 215 220
- Cys Ser Leu Tyr His Leu Leu Arg Arg Trp His Gln Arg Gly Leu Ser 225 235 235 240
- Ala Leu His Asp Phe Glu Arg Glu Tyr Thr Cys Leu Val Phe Lys Val 245 250 255
- Ser Glu Ser Arg Val Arg Phe Phe Glu His Ser Arg Val Phe Lys Asn 260 265 270
- Cys Ser Val Ala Ala Ala Pro Gly Leu Glu Leu Pro Glu Glu Gln Leu 275 280 285

His Ala Gln Val Gly Gln Ser Leu Arg Leu Phe Cys Asn Thr Ser Val Pro Ala Thr Arg Val Ala Trp Val Ser Pro Lys Asn Glu Leu Leu Val 315 Ala Pro Ala Ser Gln Asp Gly Ser Ile Ala Val Leu Ala Asp Gly Ser Leu Ala Ile Gly Arg Val Gln Glu Gln His Ala Gly Val Phe Val Cys 345 Leu Ala Ser Gly Pro Arg Leu His His Asn Gln Thr Leu Glu Tyr Asn 360 Val Ser Val Gln Lys Ala Arg Pro Glu Pro Glu Thr Phe Asn Thr Ala 370 Ala Ala Asp Pro Ile Glu Gly Arg Gly Gly Gly Gly Asp Pro Lys 395 Ser Cys Asp Lys Pro His Thr Cys Pro Pro Cys Pro Ala Pro Glu Leu Leu Gly Gly Pro Ser Val Phe Leu Phe Pro Pro Lys Pro Lys Asp Thr 425 Leu Met Ile Ser Arg Thr Pro Glu Val Thr Cys Val Val Val Asp Val Ser His Glu Asp Pro Glu Val Lys Phe Asn Trp Tyr Val Asp Gly Val Glu Val His Asn Ala Lys Thr Lys Pro Arg Glu Glu Gln Tyr Asn Ser 470 Thr Tyr Arg Val Val Ser Val Leu Thr Val Leu His Gln Asp Trp Leu 490 Asn Gly Lys Glu Tyr Lys Cys Lys Val Ser Asn Lys Ala Leu Pro Ala Pro Ile Glu Lys Thr Ile Ser Lys Ala Lys Gly Gln Pro Arg Glu Pro 520 Gln Val Tyr Thr Leu Pro Pro Ser Arg Asp Glu Leu Thr Lys Asn Gln 535 Val Ser Leu Thr Cys Leu Val Lys Gly Phe Tyr Pro Ser Asp Ile Ala Val Glu Trp Glu Ser Asn Gly Gln Pro Glu Asn Asn Tyr Lys Thr Thr Pro Pro Val Leu Asp Ser Asp Gly Ser Phe Phe Leu Tyr Ser Lys Leu Thr Val Asp Lys Ser Arg Trp Gln Gln Gly Asn Val Phe Ser Cys Ser Val Met His Glu Ala Leu His Asn His Tyr Thr Gln Lys Ser Leu Ser

610 615 620

Leu Ser Pro Gly Lys 625

<210> 13 <211> 1476 <212> DNA <213> Mus musculus <220> <221> CDS <222> (1)..(1476) <400> 13 atg caa ccc cag cgt gac ctg cga ggc ctc tgg ctc ctg ctc tcc Met Gln Pro Gln Arg Asp Leu Arg Gly Leu Trp Leu Leu Leu Ser 5 gtg ttc ctg ctt ctc ttt gag gta gcc agg gcc ggc cga tct gtg gtt Val Phe Leu Leu Phe Glu Val Ala Arg Ala Gly Arg Ser Val Val 20 agt tgt ccc gcc aac tgc ctg tgc gcc agc aac atc ctc agc tgc tcc Ser Cys Pro Ala Asn Cys Leu Cys Ala Ser Asn Ile Leu Ser Cys Ser 35 aag cag cag ctg ccc aat gtg ccc caa tct ttg ccc agc tac aca gca 192 Lys Gln Gln Leu Pro Asn Val Pro Gln Ser Leu Pro Ser Tyr Thr Ala 50 ctg ctg gac ctc agc cac aac aac ttg agc agg ctg cgg gcc gag tgg Leu Leu Asp Leu Ser His Asn Asn Leu Ser Arg Leu Arg Ala Glu Trp acc ccc acc cgg ctg acc aac ctg cac tcc ctg ctg ctg agc cac aac Thr Pro Thr Arg Leu Thr Asn Leu His Ser Leu Leu Leu Ser His Asn 85 cac ctg aac ttc atc tcc tcc gag gcc ttc gtc ccc gta ccc aac ctt 336 His Leu Asn Phe Ile Ser Ser Glu Ala Phe Val Pro Val Pro Asn Leu agg tac ttg gac ctc tcc tcc aac cat ctt cac acg ctg gat gag ttc 384 Arg Tyr Leu Asp Leu Ser Ser Asn His Leu His Thr Leu Asp Glu Phe 120 ctg ttc agc gac ctg cag gcg ctg gaa gtg ctg ttg ctc tac aat aac 432 Leu Phe Ser Asp Leu Gln Ala Leu Glu Val Leu Leu Leu Tyr Asn Asn 135 cac att gtg gtg gtg gac cgg aat gcc ttt gag gac atg gcc cag ctg His Ile Val Val Val Asp Arg Asn Ala Phe Glu Asp Met Ala Gln Leu 155 150 cag aaa ctc tac tta agc cag aat cag atc tct cgc ttt cct gtg gaa Gln Lys Leu Tyr Leu Ser Gln Asn Gln Ile Ser Arg Phe Pro Val Glu 175 165 170

ct <u>c</u> Lev	g ato	c aag e Lys	g gat S Asp 180	Gly	g aac 7 Asn	aaa Lys	tta Leu	e ccc Pro 185	Lys	a cto s Lei	g ato 1 Met	g cto Lev	ttg Lei 190	ı Asp	ctg Leu	576
tcc Ser	tco Ser	aac Asr 195	ı Lys	ctg Lev	aag Lys	aag Lys	ttg Leu 200	l Pro	cto Lev	act Thr	gac Asp	cto Leu 205	Glr	g aaa 1 Lys	ttg Leu	624
cca Pro	gco Ala 210	Trp	gtc Val	aag Lys	aat Asn	999 Gly 215	Leu	tac Tyr	ctg Lev	cat His	aac Asn 220	Asn	cco Pro	ttg Leu	gag Glu	672
tgo Cys 225	Asp	tgc Cys	aag Lys	ctc Leu	tac Tyr 230	cag Gln	ctc Leu	ttt Phe	tcg Ser	cac His	Trp	cag Gln	tac Tyr	cgg Arg	cag Gln 240	720
ctg Leu	ago Ser	tct Ser	gtg Val	atg Met 245	gac Asp	ttc Phe	cag Gln	gag Glu	gac Asp 250	Leu	tac Tyr	tgc Cys	atg Met	cac His 255	Ser	768
aag Lys	aag Lys	ctg Leu	cac His 260	aac Asn	atc Ile	ttc Phe	agc Ser	ctg Leu 265	Asp	ttc Phe	ttc Phe	aat Asn	tgc Cys 270	Ser	gag Glu	816
tac Tyr	aag Lys	gaa Glu 275	agt Ser	gcc Ala	tgg Trp	gag Glu	gct Ala 280	cac His	ctg Leu	gga Gly	gac Asp	acc Thr 285	ttg Leu	acc Thr	atc Ile	864
agg Arg	tgt Cys 290	gac Asp	acc Thr	aaa Lys	cag Gln	caa Gln 295	ggc Gly	atg Met	acc Thr	aaa Lys	gtg Val 300	tgg Trp	gtg Val	tcc Ser	cca Pro	912
agc Ser 305	aat Asn	gaa Glu	cag Gln	gtg Val	cta Leu 310	agt Ser	cag Gln	gly aaa	tcc Ser	aat Asn 315	ggc	tcg Ser	gtg Val	agc Ser	gtg Val 320	960
agg Arg	aat Asn	ggc	gac Asp	ctt Leu 325	ttt Phe	ttt Phe	aaa Lys	aag Lys	gtg Val 330	cag Gln	gtc Val	gag Glu	gat Asp	335 Gly 335	ggt Gly	1008
gtg Val	tat Tyr	acc Thr	tgt Cys 340	tac Tyr	gcc Ala	atg Met	gjà aaa	gag Glu 345	act Thr	ttc Phe	aac Asn	gag Glu	aca Thr 350	ctg Leu	tct Ser	1056
gtg Val	gag Glu	ttg Leu 355	aaa Lys	gtg Val	tat Tyr	aac Asn	ttc Phe 360	acc Thr	ttg Leu	cac His	gga Gly	cac His 365	cat His	gac Asp	acc Thr	1104
ctc Leu	aac Asn 370	aca Thr	gcc Ala	tac Tyr	act Thr	acc Thr 375	ctg Leu	gtg Val	ggc Gly	tgt Cys	atc Ile 380	ctc Leu	agt Ser	gtg Val	gtt Val	1152
ctg Leu 385	gtc Val	ctc Leu	ata Ile	tac Tyr	ttg Leu 390	tac Tyr	ctc Leu	acc Thr	cct Pro	tgc Cys 395	cgc Arg	tgc Cys	tgg Trp	tgt Cys	cgg Arg 400	1200
ggt Gly	gtg Val	gag Glu	ràs	cct Pro 405	tcc Ser	agc Ser	cac His	Gln	gga Gly 410	gat Asp	agc Ser	ctc Leu	agc Ser	tct. Ser 415	tct Ser	1248

atg Met	ctc Leu	agt Ser	acc Thr 420	aca Thr	ccc Pro	aac Asn	cac His	gac Asp 425	cct Pro	atg Met	gct Ala	ggt Gly	999 Gly 430	gac Asp	aaa Lys	1296
gat Asp	gat Asp	ggt Gly 435	ttt Phe	gac Asp	cgg Arg	cgg Arg	gtg Val 440	gcc Ala	ttc Phe	ctg Leu	gaa Glu	cct Pro 445	gct Ala	gga Gly	ccc Pro	1344
Gl <sup>A</sup> aaa	cag Gln 450	ggt Gly	caa Gln	aat Asn	ggc Gly	aaa Lys 455	atc Leu	aag Lys	cca Pro	ggc	aac Asn 460	act Thr	ctg Leu	ccg Pro	gtg Val	1392
ccc Pro 465	gaa Glu	gct Ala	aca Thr	ggc Gly	aag Lys 470	Gly	caa Gln	cgg Arg	agg Arg	atg Met 475	tcc Ser	gat Asp	cca Pro	gag Glu	tcg Ser 480	1440
gtc Val	agc Ser	tcg Ser	gtc Val	ttt Phe 485	tct Ser	gat Asp	aca Thr	ccc Pro	att Ile 490	gtg Val	gtg Val					1476

<210> 14

<211> 492

<212> PRT

<213> Mus musculus

<400> 14

Met Gln Pro Gln Arg Asp Leu Arg Gly Leu Trp Leu Leu Leu Ser 1 5 10 15

Val Phe Leu Leu Phe Glu Val Ala Arg Ala Gly Arg Ser Val Val 20 25 30

Ser Cys Pro Ala Asn Cys Leu Cys Ala Ser Asn Ile Leu Ser Cys Ser 35 40 45

Lys Gln Gln Leu Pro Asn Val Pro Gln Ser Leu Pro Ser Tyr Thr Ala 50 55 60

Leu Leu Asp Leu Ser His Asn Asn Leu Ser Arg Leu Arg Ala Glu Trp 65 70 75 80

Thr Pro Thr Arg Leu Thr Asn Leu His Ser Leu Leu Leu Ser His Asn 85 90 95

His Leu Asn Phe Ile Ser Ser Glu Ala Phe Val Pro Val Pro Asn Leu 100 105 110

Arg Tyr Leu Asp Leu Ser Ser Asn His Leu His Thr Leu Asp Glu Phe
115 120 125

Leu Phe Ser Asp Leu Gln Ala Leu Glu Val Leu Leu Leu Tyr Asn Asn 130 135 140

His Ile Val Val Val Asp Arg Asn Ala Phe Glu Asp Met Ala Gln Leu
145 150 155 160

Gln Lys Leu Tyr Leu Ser Gln Asn Gln Ile Ser Arg Phe Pro Val Glu 165 170 175

Leu Ile Lys Asp Gly Asn Lys Leu Pro Lys Leu Met Leu Leu Asp Leu

180 185 190

Ser Ser Asn Lys Leu Lys Lys Leu Pro Leu Thr Asp Leu Gln Lys Leu 195 200 205

Pro Ala Trp Val Lys Asn Gly Leu Tyr Leu His Asn Asn Pro Leu Glu 210 215 220

Cys Asp Cys Lys Leu Tyr Gln Leu Phe Ser His Trp Gln Tyr Arg Gln 225 230 235 240

Leu Ser Ser Val Met Asp Phe Gln Glu Asp Leu Tyr Cys Met His Ser 245 250 255

Lys Lys Leu His Asn Ile Phe Ser Leu Asp Phe Phe Asn Cys Ser Glu 260 265 270

Tyr Lys Glu Ser Ala Trp Glu Ala His Leu Gly Asp Thr Leu Thr Ile 275 280 285

Arg Cys Asp Thr Lys Gln Gln Gly Met Thr Lys Val Trp Val Ser Pro 290 295 300

Ser Asn Glu Gln Val Leu Ser Gln Gly Ser Asn Gly Ser Val Ser Val 305 310 315 320

Arg Asn Gly Asp Leu Phe Phe Lys Lys Val Gln Val Glu Asp Gly Gly 325 330 335

Val Tyr Thr Cys Tyr Ala Met Gly Glu Thr Phe Asn Glu Thr Leu Ser 340 345 350

Val Glu Leu Lys Val Tyr Asn Phe Thr Leu His Gly His His Asp Thr 355 360 365

Leu Asn Thr Ala Tyr Thr Thr Leu Val Gly Cys Ile Leu Ser Val Val 370 375 380

Leu Val Leu Ile Tyr Leu Tyr Leu Thr Pro Cys Arg Cys Trp Cys Arg 385 390 395 400

Gly Val Glu Lys Pro Ser Ser His Gln Gly Asp Ser Leu Ser Ser Ser 405 410 415

Met Leu Ser Thr Thr Pro Asn His Asp Pro Met Ala Gly Gly Asp Lys 420 425 430

Asp Asp Gly Phe Asp Arg Arg Val Ala Phe Leu Glu Pro Ala Gly Pro
435 440 445

Gly Gln Gly Gln Asn Gly Lys Leu Lys Pro Gly Asn Thr Leu Pro Val 450 455 460

Pro Glu Ala Thr Gly Lys Gly Gln Arg Arg Met Ser Asp Pro Glu Ser 465 470 475 480

Val Ser Ser Val Phe Ser Asp Thr Pro Ile Val Val 485 490

<210> 15 <211> 1557 <212> DNA <213> Mus musculus <220> <221> CDS <222> (1)..(1557) <400> 15 atg tcg tta agg ttc cac aca ctg ccc acc ctg cct aga gct gtc aaa Met Ser Leu Arg Phe His Thr Leu Pro Thr Leu Pro Arg Ala Val Lys 10 ccg ggt tgc aga gag ctg ctg tgt ctg ttg gtg atc gca gtg atg gtg 96 Pro Gly Cys Arg Glu Leu Leu Cys Leu Leu Val Ile Ala Val Met Val age ecc age gee tea gga atg tge ecc act get tge ate tgt gee acc 144 Ser Pro Ser Ala Ser Gly Met Cys Pro Thr Ala Cys Ile Cys Ala Thr 40 gac att gtc agc tgc acc aac aaa aac cta tct aag gtg ccc ggg aac 192 Asp Ile Val Ser Cys Thr Asn Lys Asn Leu Ser Lys Val Pro Gly Asn 55 ctt ttc aga ctg att aaa aga ctg gat ctg agc tat aac aga atc gga 240 Leu Phe Arg Leu Ile Lys Arg Leu Asp Leu Ser Tyr Asn Arg Ile Gly 65 ctg ttg gat gcc gac tgg atc ccg gtg tcg ttt gtc aag ctg agc acc 288 Leu Leu Asp Ala Asp Trp Ile Pro Val Ser Phe Val Lys Leu Ser Thr tta att ctt cgc cac aac aac atc acc agc atc tcc acg ggc agt ttc 336 Leu Ile Leu Arg His Asn Asn Ile Thr Ser Ile Ser Thr Gly Ser Phe 105 100 tcc aca acc cca aat tta aag tgt ctg gac tta tca tcc aat agg ctg 384 Ser Thr Thr Pro Asn Leu Lys Cys Leu Asp Leu Ser Ser Asn Arg Leu 120 115 aag tog gta aag agt goo aca tto caa gag otg aag got otg gaa gta Lys Ser Val Lys Ser Ala Thr Phe Gln Glu Leu Lys Ala Leu Glu Val 140 135 130 ctg ctg ctg tac aac aac cac att tcc tat ctg gac ccc gca gcg ttc 480 Leu Leu Leu Tyr Asn Asn His Ile Ser Tyr Leu Asp Pro Ala Ala Phe 160 145 150 155 ggg ggg ctt tcc cac ttg cag aaa ctc tat ctg agt ggg aac ttt ctc 528 Gly Gly Leu Ser His Leu Gln Lys Leu Tyr Leu Ser Gly Asn Phe Leu 175 165 aca cag ttc cct atg gat ttg tat act ggg agg ttc aag ctg gct gat Thr Gln Phe Pro Met Asp Leu Tyr Thr Gly Arg Phe Lys Leu Ala Asp 180 ctg aca ttt tta gat gtt tcc tat aat cga atc cct tcc ata ccg atg Leu Thr Phe Leu Asp Val Ser Tyr Asn Arg Ile Pro Ser Ile Pro Met 200

cac His	cat His 210	ata Ile	aac Asn	tta Leu	gtg Val	ccg Pro 215	gly aaa	aga Arg	cag Gln	ctg Leu	aga Arg 220	Gly	atc Ile	tac Tyr	ctt Leu	672
							gac Asp									720
							agc Ser									768
tat Tyr	acc Thr	tgt Cys	cgc Arg 260	ctg Leu	tgg Trp	tct Ser	gac Asp	tcc Ser 265	agg Arg	cac His	tcc Ser	cac His	cag Gln 270	ctg Leu	cag Gln	816
							aac Asn 280						Ile			864
							atc Ile									912
gcg Ala 305	atc Ile	gtc Val	cac His	tgt Cys	gac Asp 310	agc Ser	aag Lys	act Thr	ggc Gly	aat Asn 315	gga Gly	aat Asn	act Thr	gat Asp	ttc Phe 320	960
		_					agg Arg									1008
							aac Asn									1056
ggc	ttt Phe	gag Glu 355	gac Asp	gcc Ala	Gly 999	gta Val	tat Tyr 360	tct Ser	tgt Cys	atc Ile	gca Ala	atg Met 365	aac Asn	agg Arg	cag Gln	1104
							gat Asp									1152
							cac His									1200
acc Thr	ctg Leu	gct Ala	gcc Ala	tgc Cys 405	gtg Val	gcc Ala	agt Ser	ata Ile	gtt Val 410	cta Leu	gtg Val	cta Leu	ctg Leu	tat Tyr 415	ctg Leu	1248
							aaa Lys									1296
							cac His 440									1344

															•		
act Thi	t gg r Gl 45	· y	at g sp A	cc t la S	ct g er A	ct ga la As 45	SP AS	at co sp Ar	g as g Ly	g gc s Al	a gg .a G] 46	га гу	aa ag	ga g rg V	tc al	gtg Val	1392
tti Phe 465		g g u G	ag c lu P	cc c ro L	eu n	ag ga ys As 70	c ac	g go ır Al	g go a Al	c gg a Gl 47	y G1	ig aa In As	nt g	gc a ly L	aa ys	gtc Val 480	1440
aag Lys	g ct s Le	t ti u Pl	tc c he P	-0 0	gt g er G 85	ag ac lu Th	c gt r Va	t at	a gc e Al 49	a GI	g gg u Gl	gc at y Il	c tt e Le	eu L	ag ys 95	tcc Ser	1488
acc Thr	ag Ar	g Ai		ag t ys S 00	ct ga er As	ac tc sp Se	a ga r As	c tc p Se 50	r va	c aa l As:	t to n Se	c gt r Va	g tt l Ph 51	e Se	ca er	gac Asp	1536
aca Thr	e cc	c tt o Ph 51	ne Va	tg go	ca to la Se	c ac er Th	t ' r										1557
<21 <21	.0> : .1> : .2> 1 .3> N	519 PRT	musc	culus	3	·			•								
<40	0> 3	L6															
Met 1	Sei	Le	u Ar	g Ph	e Hi 5	s Thi	: Leu	ı Pro	Thr 10	Leu	ı Pro	o Arg	g Ala		1 1	Lys	
Pro	Gly	, Cy	s Ar 2	g G1	u Le	u Lei	ı Cys	Leu 25	Leu ;	Val	. Ile	e Ala	Va:		t v	Val	
		_	_			y Met	40	,				45	,				
Asp	Ile 50	· Va	l Se	r Cy	s Th	r Asn 55	Lys	. Asn	. Leu	Ser	Lys	val	Pro	Gl;	y P	Asn	
Leu 65	Phe	Ar	g Le	u Il	e Ly:	s Arg	Leu	Asp	Leu	Ser 75		Asn	Arg			80 80	
Leu	Leu	Ası	Al:	a As <sub>l</sub> 8!	o Trp	) Ile	Pro	Val	Ser 90	Phe	Val	Lys	Leu	Sei 95		'hr	
Leu	Ile	Let	1 Arg	g His	s Asr	Asn	Ile	Thr 105	Ser	Ile	Ser	Thr	Gly 110		e P	he	
Ser	Thr	Thr 115	Pro	Asr	ı Lev	Lys	Cys 120	Leu	Asp	Leu	Ser	Ser 125	Asn	Arc	J L	eu	
Lys	Ser 130	Val	Lys	S Ser	Ala	Thr 135	Phe	Gln	Glu	Leu	Lys 140	Ala	Leu	Glu	t Vá	al	
Leu : 145	Leu	Leu	Tyr	: Asn	Asn 150	His	Ile	Ser	Tyr	Leu 155	Asp	Pro	Ala	Ala		ne 50	
Gly (	Gly	Leu	Ser	His 165	Leu	Gln	Lys	Leu	Tyr 170	Leu	Ser	Gly	Asn	Phe 175		eu	
Thr (	Gln	Phe	Pro	Met	Asp	Leu	Tyr	Thr	Gly	Arg	Phe	Lys	Leu	Ala	As	sp.	

185 190

180 Leu Thr Phe Leu Asp Val Ser Tyr Asn Arg Ile Pro Ser Ile Pro Met 200 His His Ile Asn Leu Val Pro Gly Arg Gln Leu Arg Gly Ile Tyr Leu 215 His Gly Asn Pro Phe Val Cys Asp Cys Ser Leu Tyr Ser Leu Leu Ile Phe Trp Tyr Arg Arg His Phe Ser Ser Val Met Asp Phe Lys Asn Asp Tyr Thr Cys Arg Leu Trp Ser Asp Ser Arg His Ser His Gln Leu Gln 265 Leu Leu Gln Glu Ser Phe Leu Asn Cys Ser Tyr Ser Val Ile Asn Gly Ser Phe His Ala Leu Gly Phe Ile His Glu Ala Gln Val Gly Glu Arg Ala Ile Val His Cys Asp Ser Lys Thr Gly Asn Gly Asn Thr Asp Phe Ile Trp Val Gly Pro Asp Asn Arg Leu Leu Glu Pro Asp Lys Asp Met Gly Asn Phe Arg Val Phe Tyr Asn Gly Ser Leu Val Ile Glu Asn Pro Gly Phe Glu Asp Ala Gly Val Tyr Ser Cys Ile Ala Met Asn Arg Gln 360 Arg Leu Leu Asn Glu Thr Val Asp Ile Met Ile Asn Val Ser Asn Phe 370 Thr Ile Asn Arg Ser His Ala His Glu Ala Phe Asn Thr Ala Phe Thr Thr Leu Ala Ala Cys Val Ala Ser Ile Val Leu Val Leu Leu Tyr Leu Tyr Leu Thr Pro Cys Pro Cys Lys Cys Lys Ala Lys Arg Gln Lys Asn Thr Leu Ser Gln Ser Ser Ala His Ser Ser Ile Leu Ser Pro Gly Pro Thr Gly Asp Ala Ser Ala Asp Asp Arg Lys Ala Gly Lys Arg Val Val Phe Leu Glu Pro Leu Lys Asp Thr Ala Ala Gly Gln Asn Gly Lys Val 470 Lys Leu Phe Pro Ser Glu Thr Val Ile Ala Glu Gly Ile Leu Lys Ser Thr Arg Ala Lys Ser Asp Ser Asp Ser Val Asn Ser Val Phe Ser Asp

Thr Pro Phe Val Ala Ser Thr 515

<210> 17 <211> 1524 <212> DNA <213> Mus musculus <220> <221> CDS <222> (1)..(1524) <400> 17 atg gcc tgg cta gtg cta tca ggt ata cta cta tgc atg ttg ggt gct Met Ala Trp Leu Val Leu Ser Gly Ile Leu Leu Cys Met Leu Gly Ala 10 gga ttg ggc act tca gac ttg gag gat gtt ctg cct cct gct ccc cac Gly Leu Gly Thr Ser Asp Leu Glu Asp Val Leu Pro Pro Ala Pro His aac tgc ccc gat ata tgc atc tgt gct gcc gat gtg ttg agc tgt gcg Asn Cys Pro Asp Ile Cys Ile Cys Ala Ala Asp Val Leu Ser Cys Ala ggc cgt ggg tta cag gac ttg ccg gta gca ctg cct acc act gct gca Gly Arg Gly Leu Gln Asp Leu Pro Val Ala Leu Pro Thr Thr Ala Ala 50 gaa ctc gat ttg agc cac aac gca ctc aaa cgc ctg cac ccg ggg tgg 240 Glu Leu Asp Leu Ser His Asn Ala Leu Lys Arg Leu His Pro Gly Trp tta gcg ccc ctc tcc cgg ctg cgt gcc ttg cac cta ggc tat aat aag 288 Leu Ala Pro Leu Ser Arg Leu Arg Ala Leu His Leu Gly Tyr Asn Lys 90 ctg gaa gtc ctg ggc cat ggt gcg ttc acc aat gcc agt ggc ctg agg 336 Leu Glu Val Leu Gly His Gly Ala Phe Thr Asn Ala Ser Gly Leu Arg 105 aca ctt gac ctg tcc tct aat atg tta agg atg ctc cat acc cat gac 384 Thr Leu Asp Leu Ser Ser Asn Met Leu Arg Met Leu His Thr His Asp 115 ctg gat ggc ctg gag gag ctg gag aag tta ctt ctg ttc aat aac agc 432 Leu Asp Gly Leu Glu Glu Leu Glu Lys Leu Leu Phe Asn Asn Ser 135 ctg atg cac ttg gac ctg gat gcc ttc cag ggc ctg cgc atg ctt agc 480 Leu Met His Leu Asp Leu Asp Ala Phe Gln Gly Leu Arg Met Leu Ser 155 cac etc tat etc age tgc aac gag etc tec tet ttc tet ttc aac cac His Leu Tyr Leu Ser Cys Asn Glu Leu Ser Ser Phe Ser Phe Asn His 170 175

														•		
ttg Lei	g cac 1 His	ggt Gly	cto Lev 180	ı Gly	g tta ⁄ Leu	acc Thr	cgc	cto Leu 185	. Arg	g act g Thr	cto Lev	g gac 1 Asp	teto Lev 190	. Sea	tcc Ser	576
aac Asr	tgg Trp	cto Leu 195	1 PAS	cat His	ato Ile	tcc Ser	ato Ile 200	Pro	gag Glu	r ttg Leu	gct Ala	gca Ala 205	Let	cca Pro	a act Thr	624
tat Tyr	ctc Leu 210	Lys	aac Asn	agg Arg	cto Leu	tac Tyr 215	ctg Leu	cac His	aac Asn	aac Asn	ccg Pro	Leu	ccc Pro	tgt Cys	gac S Asp	672
tgo Cys 225	Ser	ctc Leu	tac Tyr	cac His	ctg Leu 230	Leu	cgg Arg	cgc Arg	tgg Trp	cac His 235	cag Gln	cgg Arg	ggg	cto Leu	agt Ser 240	720
gcc Ala	ctg Leu	cat His	gat Asp	ttt Phe 245	Glu	cgc Arg	gag Glu	tac Tyr	aca Thr 250	tgc Cys	ttg Leu	gtc Val	ttt Phe	aag Lys 255		768
tca Ser	gag Glu	tcc Ser	cga Arg 260	gtg Val	cgc Arg	ttt Phe	ttt Phe	gag Glu 265	cac	agc Ser	cgg Arg	gtc Val	ttc Phe 270	Lys	aac Asn	816
tgc Cys	tct Ser	gtg Val 275	gct Ala	gca Ala	gct Ala	cca Pro	ggc Gly 280	tta Leu	gag Glu	ctg Leu	cct Pro	gaa Glu 285	gag Glu	cag Gln	ctg	864
cac His	gcg Ala 290	cag Gln	gtg Val	ggc	cag Gln	tcc Ser 295	ctg Leu	agg Arg	ctc Leu	ttc Phe	tgc Cys 300	aac Asn	acc Thr	agt Ser	gtg Val	912
cct Pro 305	gcc Ala	act Thr	cgg Arg	gtg Val	gcc Ala 310	tgg Trp	gtc Val	tcc Ser	ccg Pro	aag Lys 315	aat Asn	gag Glu	ctg Leu	ctt Leu	gtg Val 320	960
gcg Ala	cca Pro	gcc Ala	tct Ser	cag Gln 325	gat Asp	ggt Gly	agc Ser	atc Ile	gct Ala 330	gtg Val	ttg Leu	gct Ala	gat Asp	ggc Gly 335	agc Ser	1008
tta Leu	gcc Ala	ata Ile	ggc Gly 340	agg Arg	gtg Val	caa Gln	gag Glu	cag Gln 345	cac His	gca Ala	ggc Gly	gtc Val	ttt Phe 350	gtg Val	tgc Cys	1056
ctg Leu	gcc Ala	agt Ser 355	ej aaa	ccc Pro	cgc Arg	ctg Leu	cac His 360	cac His	aac Asn	cag Gln	aca Thr	ctt Leu 365	gag Glu	tac Tyr	aat Asn	1104
gtg Val	agt Ser 370	gtg Val	caa Gln	aag Lys	gct Ala	cgc Arg 375	ccc Pro	gag Glu	cca Pro	gag Glu	act Thr 380	ttc Phe	aac Asn	aca Thr	ggc Gly	1152
ttt Phe 385	acc Thr	acc Thr	ctg Leu	ctg Leu	ggc ggc	tgt Cys	att Ile	gtg Val	ggc	ctg Leu 395	gtg Val	ctg Leu	gtg Val	ttg Leu	ctc Leu 400	1200
tac Tyr	ttg Leu	ttt Phe	gca Ala	cca Pro 405	ccc Pro	tgt Cys	cgt Arg	Gly	tgc Cys 410	tgt Cys	cac His	tgc Cys	tgt Cys	cag Gln· 415	cgg Arg	1248
gcc	tgc	cgc	aac	cgt	tgc	tgg	ccc	cgg	gca	tcc	agt	cca	ctc	cag	gag	1296

Ala Cys Arg Asn Arg Cys Trp Pro Arg Ala Ser Ser Pro Leu Gln Glu 420 425 ctg agc gca cag tcc tcc atg ctc agc act acg cca cca gat gca ccc Leu Ser Ala Gln Ser Ser Met Leu Ser Thr Thr Pro Pro Asp Ala Pro 435 440 age ege aag gee agt gte eae aag eat gtg gte tte etg gag eeg gge 1392 Ser Arg Lys Ala Ser Val His Lys His Val Val Phe Leu Glu Pro Gly 450 455 aag aag ggc ctc aat ggc cgt gtg cag ctc gca gta gct gaa gac ttc 1440 Lys Lys Gly Leu Asn Gly Arg Val Gln Leu Ala Val Ala Glu Asp Phe 465 470 gat ctg tgc aac ccc atg ggc ttg caa ctc aag gct ggc tct gaa tca 1488 Asp Leu Cys Asn Pro Met Gly Leu Gln Leu Lys Ala Gly Ser Glu Ser 485 490 gcc agt tcc acg ggc tca gag ggt ctc gtg atg agc 1524 Ala Ser Ser Thr Gly Ser Glu Gly Leu Val Met Ser 500 <210> 18 <211> 508 <212> PRT <213> Mus musculus <400> 18 Met Ala Trp Leu Val Leu Ser Gly Ile Leu Leu Cys Met Leu Gly Ala Gly Leu Gly Thr Ser Asp Leu Glu Asp Val Leu Pro Pro Ala Pro His 20 Asn Cys Pro Asp Ile Cys Ile Cys Ala Ala Asp Val Leu Ser Cys Ala Gly Arg Gly Leu Gln Asp Leu Pro Val Ala Leu Pro Thr Thr Ala Ala Glu Leu Asp Leu Ser His Asn Ala Leu Lys Arg Leu His Pro Gly Trp Leu Ala Pro Leu Ser Arg Leu Arg Ala Leu His Leu Gly Tyr Asn Lys Leu Glu Val Leu Gly His Gly Ala Phe Thr Asn Ala Ser Gly Leu Arg 100 105 Thr Leu Asp Leu Ser Ser Asn Met Leu Arg Met Leu His Thr His Asp 120 Leu Asp Gly Leu Glu Glu Leu Glu Lys Leu Leu Phe Asn Asn Ser

155

Leu Met His Leu Asp Leu Asp Ala Phe Gln Gly Leu Arg Met Leu Ser

His Leu Tyr Leu Ser Cys Asn Glu Leu Ser Ser Phe Ser Phe Asn His 165 170 175

- Leu His Gly Leu Gly Leu Thr Arg Leu Arg Thr Leu Asp Leu Ser Ser 180 185 190
- Asn Trp Leu Lys His Ile Ser Ile Pro Glu Leu Ala Ala Leu Pro Thr 195 200 205
- Tyr Leu Lys Asn Arg Leu Tyr Leu His Asn Asn Pro Leu Pro Cys Asp 210 215 220
- Cys Ser Leu Tyr His Leu Leu Arg Arg Trp His Gln Arg Gly Leu Ser 225 230 235 240
- Ala Leu His Asp Phe Glu Arg Glu Tyr Thr Cys Leu Val Phe Lys Val 245 250 255
- Ser Glu Ser Arg Val Arg Phe Phe Glu His Ser Arg Val Phe Lys Asn 260 265 270
- Cys Ser Val Ala Ala Ala Pro Gly Leu Glu Leu Pro Glu Glu Gln Leu 275 280 285
- His Ala Gln Val Gly Gln Ser Leu Arg Leu Phe Cys Asn Thr Ser Val 290 295 300
- Pro Ala Thr Arg Val Ala Trp Val Ser Pro Lys Asn Glu Leu Leu Val 305 310 315 320
- Ala Pro Ala Ser Gln Asp Gly Ser Ile Ala Val Leu Ala Asp Gly Ser 325 330 335
- Leu Ala Ile Gly Arg Val Gln Glu Gln His Ala Gly Val Phe Val Cys 340 345 350
- Leu Ala Ser Gly Pro Arg Leu His His Asn Gln Thr Leu Glu Tyr Asn 355 360 365
- Val Ser Val Gln Lys Ala Arg Pro Glu Pro Glu Thr Phe Asn Thr Gly 370 375 380
- Phe Thr Thr Leu Leu Gly Cys Ile Val Gly Leu Val Leu Val Leu Leu 385 390 395 400
- Tyr Leu Phe Ala Pro Pro Cys Arg Gly Cys Cys His Cys Cys Gln Arg
  405 410 415
- Ala Cys Arg Asn Arg Cys Trp Pro Arg Ala Ser Ser Pro Leu Gln Glu
  420 425 430
- Leu Ser Ala Gln Ser Ser Met Leu Ser Thr Thr Pro Pro Asp Ala Pro
  435
- Ser Arg Lys Ala Ser Val His Lys His Val Val Phe Leu Glu Pro Gly 450 455 460
- Lys Lys Gly Leu Asn Gly Arg Val Gln Leu Ala Val Ala Glu Asp. Phe 465 470 475 480
- Asp Leu Cys Asn Pro Met Gly Leu Gln Leu Lys Ala Gly Ser Glu Ser

1

485 490 495

Ala Ser Ser Thr Gly Ser Glu Gly Leu Val Met Ser 500 505

<21 <21	0 > 1 1 > 3 2 > D 3 > H	630 NA	sapi	ens												
	0> 1> C 2> (		(363	0)												
atg	0> 1: cga Arg	ccc	tcc Ser	999 999	acg Thr	gcc Ala	gly aaa	gca Ala	gcg Ala 10	ctc Leu	ctg Leu	gcg Ala	ctg Leu	ctg Leu 15	gct Ala	48
gcg Ala	ctc Leu	tgc Cys	ccg Pro 20	gcg Ala	agt Ser	cgg Arg	gct Ala	ctg Leu 25	gag Glu	gaa Glu	aag Lys	aaa Lys	gtt Val 30	tgc Cys	caa Gln	96
ggc	acg Thr	agt Ser 35	aac Asn	aag Lys	ctc Leu	acg Thr	cag Gln 40	ttg Leu	ggc Gly	act Thr	ttt Phe	gaa Glu 45	gat Asp	cat His	ttt Phe	144
ctc Leu	agc Ser 50	ctc Leu	cag Gln	agg Arg	atg Met	ttc Phe 55	aat Asn	aac Asn	tgt Cys	gag Glu	gtg Val 60	gtc Val	ctt Leu	Gly 999	aat Asn	192
ttg Leu 65	gaa Glu	att Ile	acc Thr	tat Tyr	gtg Val 70	cag Gln	agg Arg	aat Asn	tat Tyr	gat Asp 75	ctt Leu	tcc Ser	ttc Phe	tta Leu	aag Lys 80	240
acc Thr	atc Ile	cag Gln	gag Glu	gtg Val 85	gct Ala	ggt Gly	tat Tyr	gtc Val	ctc Leu 90	att Ile	gcc Ala	ctc Leu	aac Asn	aca Thr 95	gtg Val	288
gag Glu	cga Arg	att Ile	cct Pro 100	ttg Leu	gaa Glu	aac Asn	ctg Leu	cag Gln 105	atc Ile	atc Ile	aga Arg	gga Gly	aat Asn 110	atg Met	tac Tyr	336
tac Tyr	gaa Glu	aat Asn 115	tcc Ser	tat Tyr	gcc Ala	tta Leu	gca Ala 120	gtc Val	tta Leu	tct Ser	aac Asn	tat Tyr 125	gat Asp	gca Ala	aat Asn	384
aaa Lys	acc Thr 130	gga Gly	ctg Leu	aag Lys	gag Glu	ctg Leu 135	ccc Pro	atg Met	aga Arg	aat Asn	tta Leu 140	cag Gln	gaa Glu	atc Ile	ctg Leu	432
cat His 145	ggc	gcc Ala	gtg Val	cgg Arg	ttc Phe 150	agc Ser	aac Asn	aac Asn	cct Pro	gcc Ala 155	ctg Leu	tgc Cys	aac Asn	gtg Val	gag Glu 160	480
agc Ser	atc Ile	cag Gln	tgg Trp	cgg Arg 165	gac Asp	ata Ile	gtc Val	agc Ser	agt Ser 170	gac Asp	ttt Phe	ctc Leu	agc Ser	aac Asn 175	atg Met	528

tcg Ser	g ato	gad : Asp	tto Phe 180	Glr	g aac 1 Asn	cac His	cto Lei	9 990 1 Gly 185	Ser	tgo Cys	caa Glr	a aag 1 Lys	g tgt s Cys 190	s Asj	cca Pro	576
ago Ser	tgt Cys	ccc Pro 195	Asr.	ggs Gly	g ago Ser	tgc Cys	tgg Trp 200	Gly	gca Ala	ı gga	gag Glu	g gag ı Glu 205	ı Asr	tgo LCys	c cag s Gln	624
aaa Lys	cto Leu 210	Thr	aaa Lys	ato Ile	ato Ile	tgt Cys 215	Ala	cag Gln	cag Glm	tgo Cys	s Ser 220	: Gly	g cgc Arg	tgo g Cys	c cgt 3 Arg	672
ggc Gly 225	Lys	tcc Ser	e ccc	agt Ser	gac Asp 230	Cys	tgo Cys	cac His	aac Asn	Gln 235	Cys	gct Ala	gca Ala	ggo Gly	tgc Cys 240	720
aca Thr	. Gly	e ccc	cgg Arg	gag Glu 245	Ser	gac Asp	tgc Cys	ctg Leu	gtc Val 250	Cys	cgc Arc	: aaa   Lys	ttc Phe	cga Arg 255	a gac J.Asp	768
gaa Glu	gcc Ala	acg Thr	tgc Cys 260	Lys	gac Asp	acc Thr	tgc Cys	Pro 265	cca Pro	cto Leu	atg Met	ctc Leu	tac Tyr 270	Asr	ccc Pro	816
acc Thr	acg Thr	tac Tyr 275	GIn	atg Met	gat Asp	gtg Val	aac Asn 280	Pro	gag Glu	ggc Gly	aaa Lys	tac Tyr 285	Ser	ttt Phe	ggt Gly	864
gcc Ala	acc Thr 290	tgc Cys	gtg Val	aag Lys	aag Lys	tgt Cys 295	ccc Pro	cgt Arg	aat Asn	tat Tyr	gtg Val 300	gtg Val	aca Thr	gat Asp	cac	912
ggc 305	tcg Ser	tgc Cys	gtc Val	cga Arg	gcc Ala 310	tgt Cys	GJÀ aaa	gcc Ala	gac Asp	agc Ser 315	tat Tyr	gag Glu	atg Met	gag Glu	gaa Glu 320	960
gac Asp	ggc	gtc Val	cgc Arg	aag Lys 325	tgt Cys	aag Lys	aag Lys	tgc Cys	gaa Glu 330	Gly	cct Pro	tgc Cys	cgc Arg	aaa Lys 335	gtg Val	1008
tgt Cys	aac Asn	gga Gly	ata Ile 340	ggt Gly	att Ile	ggt Gly	gaa Glu	ttt Phe 345	aaa Lys	gac Asp	tca Ser	ctc Leu	tcc Ser 350	ata Ile	aat Asn	1056
gct Ala	acg Thr	aat Asn 355	att Ile	aaa Lys	cac His	ttc Phe	aaa Lys 360	aac Asn	tgc Cys	acc Thr	tcc Ser	atc Ile 365	agt Ser	Gly	gat Asp	1104
ctc Leu	cac His 370	atc Ile	ctg Leu	ccg Pro	gtg Val	gca Ala 375	ttt Phe	agg Arg	ggt Gly	gac Asp	tcc Ser 380	ttc Phe	aca Thr	cat His	act Thr	1152
cct Pro 385	cct Pro	ctg Leu	gat Asp	cca Pro	cag Gln 390	gaa Glu	ctg Leu	gat Asp	att Ile	ctg Leu 395	aaa Lys	acc Thr	gta Val	aag Lys	gaa Glu 400	1200
atc Ile	aca Thr	Gly 999	ttt Phe	ttg Leu 405	ctg Leu	att Ile	cag Gln	gct Ala	tgg Trp 410	cct Pro	gaa Glu	aac Asn	agg Arg	acg Thr 415	gac Asp	1248
ctc	cat	gcc	ttt	gag	aac	cta	gaa	atc	ata	cgc	ggc	agg	acc	aag	caa	1296

Leu His A	Ala Phe 420	Glu A	Asn L	eu (		Ile 425	Ile	Arg	Gly	Arg	Thr 430	Lys	Gln	
cat ggt c His Gly G 4	cag ttt 31n Phe 135	tct c Ser I	ctt g Leu A	la V	gtc Val 440	gtc Val	agc Ser	ctg Leu	aac Asn	ata Ile 445	aca Thr	tcc Ser	ttg Leu	1344
gga tta c Gly Leu A 450			Lys G											1392
gga aac a Gly Asn I 465		Leu C												1440
ttt ggg a Phe Gly T													Glu	1488
aac agc t Asn Ser C					Gln									1536
gag ggc t Glu Gly C				lu 1			_	_	_		-			1584
gtc agc c Val Ser A 530			Glu C											1632
gag cca a Glu Pro A 545		Phe V												1680
gag tgc c Glu Cys I														1728
gac aac t Asp Asn C														1776
aag acc t Lys Thr (	tgc ccg Cys Pro 595	gca g Ala (	gga g Gly V	/al I	atg Met 600	gga Gly	gaa Glu	aac Asn	aac Asn	acc Thr 605	ctg Leu	gtc Val	tgg Trp	1824
aag tac g Lys Tyr A 610			Gly H											1872
acc tac of Thr Tyr 0 625		Thr (												1920
cct aag a Pro Lys 1														1968

660 665 670

ato Ile	gtt Val	cgg Arg 675	, Гуз	g cgo : Arg	aco Thr	cto Lev	g cgg Arg 680	J Arg	g cto J Lei	g cto 1 Lei	g caq ı Glr	g gag n Glu 685	ı Arg	g gag g Glu	g ctt 1 Leu	2064
gtg Val	gag Glu	Pro	ctt Leu	aca Thr	ccc Pro	s agt Ser 695	Gly	gaa Glu	gct Ala	cco Pro	aac Asr 700	ı Gln	gct Ala	cto Lev	ttg Leu	2112
agg Arg 705	ITe	ttg Leu	aag Lys	gaa Glu	act Thr 710	Glu	ttc Phe	aaa Lys	aag Lys	ato Ile 715	Lys	gtg Val	ctg Leu	ggo	tcc Ser 720	2160
ggt Gly	gcg Ala	ttc Phe	ggc	acg Thr 725	Val	tat Tyr	aag Lys	gga Gly	Leu 730	Trp	ato Ile	cca Pro	gaa Glu	ggt Gly 735	gag Glu	2208
aaa Lys	gtt Val	aaa Lys	att Ile 740	ccc Pro	gtc Val	gct Ala	atc Ile	aag Lys 745	gaa Glu	tta Leu	aga Arg	gaa Glu	gca Ala 750	aca Thr	tct Ser	2256
ccg Pro	aaa Lys	gcc Ala 755	aac Asn	aag Lys	gaa Glu	atc Ile	ctc Leu 760	gat Asp	gaa Glu	gcc Ala	tac Tyr	gtg Val 765	atg Met	gcc Ala	agc Ser	2304
gtg Val	gac Asp 770	aac Asn	ccc Pro	cac	gtg Val	tgc Cys 775	cgc Arg	ctg Leu	ctg Leu	Gly	atc Ile 780	tgc Cys	ctc Leu	acc Thr	tcc Ser	2352
acc Thr 785	gtg Val	cag Gln	ctc Leu	atc Ile	acg Thr 790	cag Gln	ctc Leu	atg Met	ccc Pro	ttc Phe 795	Gly	tgc Cys	ctc Leu	ctg Leu	gac Asp 800	2400
tat Tyr	gtc Val	cgg Arg	gaa Glu	cac His 805	aaa Lys	gac Asp	aat Asn	att Ile	ggc Gly 810	tcc Ser	cag Gln	tac Tyr	ctg Leu	ctc Leu 815	aac Asn	2448
tgg Trp	tgt Cys	gtg Val	cag Gln 820	atc Ile	gca Ala	aag Lys	ggc Gly	atg Met 825	aac Asn	tac Tyr	ttg Leu	gag Glu	gac Asp 830	cgt Arg	cgc Arg	2496
ttg Leu	gtg Val	cac His 835	cgc Arg	gac Asp	ctg Leu	gca Ala	gcc Ala 840	agg Arg	aac Asn	gta Val	ctg Leu	gtg Val 845	aaa Lys	aca Thr	ccg Pro	2544
cag Gln	cat His 850	gtc Val	aag Lys	atc Ile	aca Thr	gat Asp 855	ttt Phe	G1y 999	ctg Leu	gcc Ala	aaa Lys 860	ctg Leu	ctg Leu	ggt Gly	gcg Ala	2592
gaa Glu 865	gag Glu	aaa Lys	gaa Glu	tac Tyr	cat His 870	gca Ala	gaa Glu	gga Gly	ggc Gly	aaa Lys 875	gtg Val	cct Pro	atc Ile	aag Lys	tgg Trp 880	2640
atg Met	gca Ala	ttg Leu	Glu	tca Ser 885	att Ile	tta Leu	cac His	aga Arg	atc Ile 890	tat Tyr	acc Thr	cac His	Gln	agt Ser 895	gat Asp	2688
gtc Val	tgg Trp	Ser	tac Tyr 900	Gly aaa	gtg Val	acc Thr	Val	tgg Trp 905	gag Glu	ttg Leu	atg Met	Thr	ttt Phe 910	gga Gly	tcc Ser	2736

aag Lys	g cca	a tat Tyr 915	Asp	gga	ato Ile	cct Pro	gco Ala 920	Ser	gag Glu	g ato	tcc Ser	Ser 925	Ile	cto Leu	g gag ı Glu	2784
aaa Lys	930 930	GIU	cgc Arg	ctc Leu	cct Pro	cag Glr 935	Pro	cco Pro	ata Ile	tgt Cys	acc Thr 940	Ile	gat Asp	gto Val	tac Tyr	2832
ats Met 945	: TŤ€	atg Met	gtc Val	aag Lys	tgc Cys 950	Trp	atg Met	ata Ile	gac Asp	gca Ala 955	Asp	agt Ser	cgc Arg	cca Pro	aag Lys 960	2880
ttc Phe	cgt Arg	gag Glu	ttg Leu	atc Ile 965	Ile	gaa Glu	ttc Phe	tcc Ser	aaa Lys 970	Met	gcc Ala	cga Arg	gac Asp	ccc Pro 975	Gln	2928
cgc Arg	tac Tyr	ctt Leu	gtc Val 980	att Ile	cag Gln	gly aaa	gat Asp	gaa Glu 985	aga Arg	atg Met	cat His	ttg Leu	cca Pro 990	agt Ser	cct Pro	2976
aca Thr	. gac	Ser 995	aac Asn	ttc Phe	tac Tyr	Arg	gcc Ala 1000	ctg Leu	atg Met	gat Asp	gaa Glu	gaa Glu 1005	gac Asp	atg Met	gac Asp	3024
Asp	gtg Val 1010	Val	gat Asp	gcc Ala	Asp	gag Glu 1015	tac Tyr	ctc Leu	atc Ile	Pro	cag Gln 1020	cag Gln	ggc Gly	ttc Phe	ttc Phe	3072
agc Ser 102	ser	ccc Pro	tcc Ser	Thr	tca Ser 1030	cgg Arg	act Thr	ccc Pro	Leu	ctg Leu 1035	agc Ser	tct Ser	ctg Leu	Ser	gca Ala 1040	3120
THE	ser	Asn	Asn 1	Ser .045	Thr	Val	Ala	Cys 1	Ile LOSO	Asp	aga Arg	Asn	Gly	Leu L055	Gln	3168
per	Cys	Pro	11e	гÀг	Glu	Asp	Ser 1	Phe 1065	Leu	Gln	cga Arg	Tyr 1	Ser .070	Ser	Asp	3216
PIO	inr	G1y 1075	Ala	Leu	Thr	Glu 1	qaA 080.	Ser	Ile	Asp		Thr 085	Phe	Leu	Pro	3264
vai 1	1090	GIU	Tyr	TTE	Asn 1	G1n 095	Ser	Val	Pro	Lys 1	agg Arg .100	Pro	Ala	Gly	Ser	3312
1105	GIN	ASN	Pro	vaı 1	Tyr 110	His	Asn	Gln	Pro 1	Leu 115	aac Asn	Pro .	Ala	Pro 1	Ser 120	3360
arg	Asp	Pro	ніs : 1:	lyr 125	GIn .	Asp	Pro	His 1	Ser 130	Thr	gca q Ala	Val (	Gly .	Asn 135	Pro	3408
gag Glu	tat Tyr	ren .	aac a Asn : 140	act Thr	gtc ( Val (	cag Gln	Pro '	acc Thr (	tgt Cys	gtc Val	aac a Asn s	Ser :	aca Thr 1	ttc. Phe	gac Asp	3456

agc Ser	Pro	gcc Ala 1155	His	tgg Trp	gcc Ala	Gln	aaa Lys 1160	Gly	agc Ser	cac His	Gln	att Ile 1165	agc Ser	ctg Leu	gac Asp	3504
Asn	cct Pro 1170	gac Asp	tac Tyr	cag Gln	Gln	gac Asp 1175	ttc Phe	ttt Phe	ccc Pro	Lys	gaa Glu 1180	gcc Ala	aag Lys	cca Pro	aat Asn	3552
ggc Gly 118!	Ile	ttt Phe	aag Lys	Gly	tcc Ser 1190	aca Thr	gct Ala	gaa Glu	Asn	gca Ala 1195	gaa Glu	tac Tyr	cta Leu	agg Arg	gtc Val 1200	3600
gcg Ala	cca Pro	caa Gln	Ser	agt Ser 1205	gaa Glu	ttt Phe	att Ile	Gly	gca Ala 1210							3630
<213	0 > 2 : 1 > 1 : 2 > P!	210										:				
<213	3 > H	omo	sapi	ens									-			
	0> 20					•										
Met 1	Arg	Pro	Ser	Gly 5	Thr	Ala	Gly	Ala	Ala 10	Leu	Leu	Ala	Leu	Leu 15	Ala	
Ala	Leu	Cys	Pro 20	Ala	Ser	Arg	Ala	Leu 25	Glu	Glu	ГЛЗ	Lys	Val 30	Cys	Gln	
Gly	Thr	Ser 35	Asn	Lys	Leu	Thr	Gln 40	Leu	Gly	Thr	Phe	Glu 45	Asp	His	Phe	
Leu	Ser 50	Leu	Gln	Arg	Met	Phe 55	Asn	Asn	Cys	Glu	Val 60	Val	Leu	Gly	Asn	
Leu 65	Glu	Ile	Thr	Tyr	Val 70	Gln	Arg	Asn	Tyr	Asp 75	Leu	Ser	Phe	Leu	Lys 80	
Thr	Ile	Ġln	Glu	Val 85	Ala	Gly	Tyr	Val	Leu 90	Ile	Ala	Leu	Asn	Thr 95	Val	
Glu	Arg	Ile	Pro 100	Leu	Glu	Asn	Leu	Gln 105	Ile	Ile	Arg	Gly	Asn 110	Met	Tyr	
Tyr	Glu	Asn 115	Ser	Tyr	Ala	Leu	Ala 120	Val	Leu	Ser	Asn	Tyr 125	Asp	Ala	Asn	
Lys	Thr 130	Gly	Leu	Lys	Glu	Leu 135	Pro	Met	Arg	Asn	Leu 140	Gln	Glu	Ile	Leu	
His 145	Gly	Ala	Val	Arg	Phe 150	Ser	Asn	Asn	Pro	Ala 155	Leu	Cys	Asn	Val	Glu 160	
Ser	Ile	Gln	Trp	Arg 165	Asp	Ile	Val	Ser	Ser 170	Asp	Phe	Leu	Ser	Asn 175	Met	
Ser	Met	Asp	Phe 180	Gln	Asn	His	Leu	Gly 185	Ser	Cys	Gln	Lys	Cys 190	Asp	Pro	
Ser	Cys	Pro	Asn	Gly	Ser	Cys	Trp	Gly	Ala	Gly	Glu	Glu	Asn	Cys	Gln	

195 200 205

Lys Leu Thr Lys Ile Ile Cys Ala Gln Gln Cys Ser Gly Arg Cys Arg 210 215 220

Gly Lys Ser Pro Ser Asp Cys Cys His Asn Gln Cys Ala Ala Gly Cys 225 230 235 240

Thr Gly Pro Arg Glu Ser Asp Cys Leu Val Cys Arg Lys Phe Arg Asp 245 250 255

Glu Ala Thr Cys Lys Asp Thr Cys Pro Pro Leu Met Leu Tyr Asn Pro 260 265 270

Thr Thr Tyr Gln Met Asp Val Asn Pro Glu Gly Lys Tyr Ser Phe Gly 275 280 285

Ala Thr Cys Val Lys Lys Cys Pro Arg Asn Tyr Val Val Thr Asp His 290 295 300

Gly Ser Cys Val Arg Ala Cys Gly Ala Asp Ser Tyr Glu Met Glu Glu 305 310 315 320

Asp Gly Val Arg Lys Cys Lys Lys Cys Glu Gly Pro Cys Arg Lys Val 325 330 335

Cys Asn Gly Ile Gly Ile Gly Glu Phe Lys Asp Ser Leu Ser Ile Asn 340 345 350

Ala Thr Asn Ile Lys His Phe Lys Asn Cys Thr Ser Ile Ser Gly Asp 355 360 365

Leu His Ile Leu Pro Val Ala Phe Arg Gly Asp Ser Phe Thr His Thr 370 375 380

Pro Pro Leu Asp Pro Gln Glu Leu Asp Ile Leu Lys Thr Val Lys Glu 385 390 395 400

Ile Thr Gly Phe Leu Leu Ile Gln Ala Trp Pro Glu Asn Arg Thr Asp 405 410 415

Leu His Ala Phe Glu Asn Leu Glu Ile Ile Arg Gly Arg Thr Lys Gln
420 425 430

His Gly Gln Phe Ser Leu Ala Val Val Ser Leu Asn Ile Thr Ser Leu 435 440 445

Gly Leu Arg Ser Leu Lys Glu Ile Ser Asp Gly Asp Val Ile Ile Ser 450 455 460

Gly Asn Lys Asn Leu Cys Tyr Ala Asn Thr Ile Asn Trp Lys Lys Leu 465 470 475 480

Phe Gly Thr Ser Gly Gln Lys Thr Lys Ile Ile Ser Asn Arg Gly Glu 485 490 495

Asn Ser Cys Lys Ala Thr Gly Gln Val Cys His Ala Leu Cys Ser Pro 500 505 510

Glu Gly Cys Trp Gly Pro Glu Pro Arg Asp Cys Val Ser Cys Arg Asn

515 520 525

Val Ser Arg Gly Arg Glu Cys Val Asp Lys Cys Asn Leu Leu Glu Gly 530 535 540

Glu Pro Arg Glu Phe Val Glu Asn Ser Glu Cys Ile Gln Cys His Pro 545 550 555 560

Glu Cys Leu Pro Gln Ala Met Asn Ile Thr Cys Thr Gly Arg Gly Pro 565 570 575

Asp Asn Cys Ile Gln Cys Ala His Tyr Ile Asp Gly Pro His Cys Val 580 585 590

Lys Thr Cys Pro Ala Gly Val Met Gly Glu Asn Asn Thr Leu Val Trp 595 600 605

Lys Tyr Ala Asp Ala Gly His Val Cys His Leu Cys His Pro Asn Cys 610 620

Thr Tyr Gly Cys Thr Gly Pro Gly Leu Glu Gly Cys Pro Thr Asn Gly 625 635 635

Pro Lys Ile Pro Ser Ile Ala Thr Gly Met Val Gly Ala Leu Leu Leu 645 650 655

Leu Leu Val Val Ala Leu Gly Ile Gly Leu Phe Met Arg Arg Arg His 660 665 670

Ile Val Arg Lys Arg Thr Leu Arg Arg Leu Leu Gln Glu Arg Glu Leu 675 680 685

Val Glu Pro Leu Thr Pro Ser Gly Glu Ala Pro Asn Gln Ala Leu Leu 690 695 700

Arg Ile Leu Lys Glu Thr Glu Phe Lys Lys Ile Lys Val Leu Gly Ser 705 710 715 720

Gly Ala Phe Gly Thr Val Tyr Lys Gly Leu Trp Ile Pro Glu Gly Glu
725 730 735

Lys Val Lys Ile Pro Val Ala Ile Lys Glu Leu Arg Glu Ala Thr Ser

Pro Lys Ala Asn Lys Glu Ile Leu Asp Glu Ala Tyr Val Met Ala Ser

Val Asp Asn Pro His Val Cys Arg Leu Leu Gly Ile Cys Leu Thr Ser 770 775 780

Thr Val Gln Leu Ile Thr Gln Leu Met Pro Phe Gly Cys Leu Leu Asp 785 790 795 800

Tyr Val Arg Glu His Lys Asp Asn Ile Gly Ser Gln Tyr Leu Leu Asn 805 810 815

Trp Cys Val Gln Ile Ala Lys Gly Met Asn Tyr Leu Glu Asp Arg Arg 820 825 830 .

Leu Val His Arg Asp Leu Ala Ala Arg Asn Val Leu Val Lys Thr Pro 835 840 845

1

Gln His Val Lys Ile Thr Asp Phe Gly Leu Ala Lys Leu Leu Gly Ala 850 855

- Glu Glu Lys Glu Tyr His Ala Glu Gly Gly Lys Val Pro Ile Lys Trp 865 870 875 880
- Met Ala Leu Glu Ser Ile Leu His Arg Ile Tyr Thr His Gln Ser Asp 885 890 895
- Val Trp Ser Tyr Gly Val Thr Val Trp Glu Leu Met Thr Phe Gly Ser 900 905 910
- Lys Pro Tyr Asp Gly Ile Pro Ala Ser Glu Ile Ser Ser Ile Leu Glu 915 920 925
- Lys Gly Glu Arg Leu Pro Gln Pro Pro Ile Cys Thr Ile Asp Val Tyr 930 935 940
- Met Ile Met Val Lys Cys Trp Met Ile Asp Ala Asp Ser Arg Pro Lys 945 950 955 960
- Phe Arg Glu Leu Ile Ile Glu Phe Ser Lys Met Ala Arg Asp Pro Gln 965 970 975
- Arg Tyr Leu Val Ile Gln Gly Asp Glu Arg Met His Leu Pro Ser Pro 980 985 990
- Thr Asp Ser Asn Phe Tyr Arg Ala Leu Met Asp Glu Glu Asp Met Asp 995 1000 1005
- Asp Val Val Asp Ala Asp Glu Tyr Leu Ile Pro Gln Gln Gly Phe Phe 1010 1015 1020
- Ser Ser Pro Ser Thr Ser Arg Thr Pro Leu Leu Ser Ser Leu Ser Ala 025 1030 1035 1040
- Thr Ser Asn Asn Ser Thr Val Ala Cys Ile Asp Arg Asn Gly Leu Gln
  1045 1050 1055
- Ser Cys Pro Ile Lys Glu Asp Ser Phe Leu Gln Arg Tyr Ser Ser Asp 1060 1065 1070
- Pro Thr Gly Ala Leu Thr Glu Asp Ser Ile Asp Asp Thr Phe Leu Pro 1075 1080 1085
- Val Pro Glu Tyr Ile Asn Gln Ser Val Pro Lys Arg Pro Ala Gly Ser 1090 1095 1100
- Val Gln Asn Pro Val Tyr His Asn Gln Pro Leu Asn Pro Ala Pro Ser 105 1110 1115 1120
- Arg Asp Pro His Tyr Gln Asp Pro His Ser Thr Ala Val Gly Asn Pro 1125 1130 1135
- Glu Tyr Leu Asn Thr Val Gln Pro Thr Cys Val Asn Ser Thr Phe Asp 1140 1145 1150
- Ser Pro Ala His Trp Ala Gln Lys Gly Ser His Gln Ile Ser Leu Asp 1155 1160 1165

Asn Pro Asp Tyr Gln Gln Asp Phe Phe Pro Lys Glu Ala Lys Pro Asn 1175 Gly Ile Phe Lys Gly Ser Thr Ala Glu Asn Ala Glu Tyr Leu Arg Val 1190 1195 Ala Pro Gln Ser Ser Glu Phe Ile Gly Ala 1205 <210> 21 <211> 5512 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (167)..(3799) <220> <223> Human EGFR <400> 21 gagetagece eggeggeege egeegeeeag aceggaegae aggeeacete gteggegtee 60 geocgagtee eegectegee gecaacgeea caaccacege geacggeeee etgacteegt 120 ccagtattga tegggagage eggagegage tettegggga geageg atg ega eec Met Arg Pro tec ggg acg gcc ggg gca gcg ctc ctg gcg ctg ctg gct gcg ctc tgc 223 Ser Gly Thr Ala Gly Ala Ala Leu Leu Ala Leu Leu Ala Ala Leu Cys ccg gcg agt cgg gct ctg gag gaa aag aaa gtt tgc caa ggc acg agt 271 Pro Ala Ser Arg Ala Leu Glu Glu Lys Lys Val Cys Gln Gly Thr Ser aac aag ctc acg cag ttg ggc act ttt gaa gat cat ttt ctc agc ctc 319 Asn Lys Leu Thr Gln Leu Gly Thr Phe Glu Asp His Phe Leu Ser Leu cag agg atg ttc aat aac tgt gag gtg gtc ctt ggg aat ttg gaa att Gln Arg Met Phe Asn Asn Cys Glu Val Val Leu Gly Asn Leu Glu Ile acc tat gtg cag agg aat tat gat ctt tcc ttc tta aag acc atc cag 415 Thr Tyr Val Gln Arg Asn Tyr Asp Leu Ser Phe Leu Lys Thr Ile Gln gag gtg gct ggt tat gtc ctc att gcc ctc aac aca gtg gag cga att Glu Val Ala Gly Tyr Val Leu Ile Ala Leu Asn Thr Val Glu Arg Ile cct ttg gaa aac ctg cag atc atc aga gga aat atg tac tac gaa aat 511 Pro Leu Glu Asn Leu Gln Ile Ile Arg Gly Asn Met Tyr Tyr Glu Asn 100 105 · 11:5 tcc tat gcc tta gca gtc tta tct aac tat gat gca aat aaa acc gga Ser Tyr Ala Leu Ala Val Leu Ser Asn Tyr Asp Ala Asn Lys Thr Gly

125 120 130 ctg aag gag ctg ccc atg aga aat tta cag gaa atc ctg cat ggc gcc 607 Leu Lys Glu Leu Pro Met Arg Asn Leu Gln Glu Ile Leu His Gly Ala gtg cgg ttc agc aac acc cct gcc ctg tgc aac gtg gag agc atc cag Val Arg Phe Ser Asn Asn Pro Ala Leu Cys Asn Val Glu Ser Ile Gln tgg cgg gac ata gtc agc agt gac ttt ctc agc aac atg tcg atg gac 703 Trp Arg Asp Ile Val Ser Ser Asp Phe Leu Ser Asn Met Ser Met Asp tto cag aac cac ctg ggc agc tgc caa aag tgt gat cca agc tgt ccc 751 Phe Gln Asn His Leu Gly Ser Cys Gln Lys Cys Asp Pro Ser Cys Pro aat ggg agc tgc tgg ggt gca gga gag gag aac tgc cag aaa ctg acc 799 Asn Gly Ser Cys Trp Gly Ala Gly Glu Glu Asn Cys Gln Lys Leu Thr aaa atc atc tgt gcc cag cag tgc tcc ggg cgc tgc cgt ggc aag tcc 847 Lys Ile Ile Cys Ala Gln Gln Cys Ser Gly Arg Cys Arg Gly Lys Ser ccc agt gac tgc tgc cac aac cag tgt gct gca ggc tgc aca ggc ccc 895 Pro Ser Asp Cys Cys His Asn Gln Cys Ala Ala Gly Cys Thr Gly Pro 230 cgg gag agc gac tgc ctg gtc tgc cgc aaa ttc cga gac gaa gcc acg 943 Arg Glu Ser Asp Cys Leu Val Cys Arg Lys Phe Arg Asp Glu Ala Thr tgc aag gac acc tgc ccc cca ctc atg ctc tac aac ccc acc acg tac 991 Cys Lys Asp Thr Cys Pro Pro Leu Met Leu Tyr Asn Pro Thr Tyr 260 270 cag atg gat gtg aac ccc gag ggc aaa tac agc ttt ggt gcc acc tqc 1039 Gln Met Asp Val Asn Pro Glu Gly Lys Tyr Ser Phe Gly Ala Thr Cys 280 gtg aag aag tgt ccc cgt aat tat gtg gtg aca gat cac ggc tcg tgc 1087 Val Lys Lys Cys Pro Arg Asn Tyr Val Val Thr Asp His Gly Ser Cys 300 gtc cga gcc tgt ggg gcc gac agc tat gag atg gag gaa gac ggc gtc 1135 Val Arg Ala Cys Gly Ala Asp Ser Tyr Glu Met Glu Glu Asp Gly Val 310 cgc aag tgt aag aag tgc gaa ggg cct tgc cgc aaa gtg tgt aac gga 1183 Arg Lys Cys Lys Cys Glu Gly Pro Cys Arg Lys Val Cys Asn Gly 330 ata ggt att ggt gaa ttt aaa gac tca ctc tcc ata aat gct acg aat 1231 Ile Gly Ile Gly Glu Phe Lys Asp Ser Leu Ser Ile Asn Ala Thr Asn 340 345 350 355 att aaa cac ttc aaa aac tgc acc tcc atc agt ggc gat ctc cac atc Ile Lys His Phe Lys Asn Cys Thr Ser Ile Ser Gly Asp Leu His Ile 360

ctg Leu	ccg Pro	gtg Val	gca Ala 375	Phe	agg Arg	ggt	gac Asp	Ser 380	. Phe	aca Thr	cat His	act Thr	cct Pro 385	Pro	ctg Leu	1327
gat Asp	cca Pro	cag Gln 390	Glu	ctg Leu	gat Asp	att Ile	ctg Leu 395	. Lys	acc Thr	gta Val	aag . Lys	gaa Glu 400	Ile	aca Thr	Gly 999	1375
ttt Phe	ttg Leu 405	Leu	att Ile	cag Gln	gct Ala	tgg Trp 410	Pro	gaa Glu	aac Asn	agg Arg	acg Thr 415	Asp	cto Leu	cat His	gcc Ala	1423
ttt Phe 420	Glu	aac Asn	cta Leu	gaa Glu	atc Ile 425	ata Ile	cgc Arg	ggc	agg Arg	acc Thr 430	Lys	caa Gln	cat His	ggt Gly	cag Gln 435	1471
ttt Phe	tct Ser	ctt Leu	gca Ala	gtc Val 440	Val	agc Ser	ctg Leu	aac Asn	ata Ile 445	Thr	tcc Ser	ttg Leu	gga Gly	Leu 450	cgc Arg	1519
tcc Ser	ctc Leu	aag Lys	gag Glu 455	ata Ile	agt Ser	gat Asp	gga Gly	gat Asp 460	gtg Val	ata Ile	att Ile	tca Ser	gga Gly 465	aac Asn	aaa Lys	1567
aat Asn	ttg Leu	tgc Cys 470	tat Tyr	gca Ala	aat Asn	aca Thr	ata Ile 475	aac Asn	tgg Trp	aaa Lys	aaa Lys	ctg Leu 480	ttt Phe	gly aaa	acc Thr	1615
tcc Ser	ggt Gly 485	cag Gln	aaa Lys	acc Thr	aaa Lys	att Ile 490	ata Ile	agc Ser	aac Asn	aga Arg	ggt Gly 495	gaa Glu	aac Asn	agc Ser	tgc Cys	1663
aag Lys 500	gcc Ala	aca Thr	Gly	cag Gln	gtc Val 505	tgc Cys	cat His	gcc Ala	ttg Leu	tgc Cys 510	tcc Ser	ccc Pro	gag Glu	ggc	tgc Cys 515	1711 ·
tgg Trp	ggc	ccg Pro	gag Glu	ccc Pro 520	agg Arg	gac Asp	tgc Cys	gtc Val	tct Ser 525	tgc Cys	cgg Arg	aat Asn	gtc Val	agc Ser 530	cga Arg	1759
ggc Gly	agg Arg	gaa Glu	tgc Cys 535	gtg Val	gac Asp	aag Lys	tgc Cys	aaç Asn 540	ctt Leu	ctg Leu	gag Glu	ggt Gly	gag Glu 545	cca Pro	agg Arg	1807
gag Glu	ttt Phe	gtg Val 550	gag Glu	aac Asn	tct Ser	gag Glu	tgc Cys 555	ata Ile	cag Gln	tgc Cys	cac His	cca Pro 560	gag Glu	tgc Cys	ctg Leu	1855
cct Pro	cag Gln 565	gcc Ala	atg Met	aac Asn	atc Ile	acc Thr 570	tgc Cys	aca Thr	gga Gly	cgg Arg	gga Gly 575	cca Pro	gac Asp	aac Asn	tgt Cys	1903
atc Ile 580	cag Gln	tgt Cys	gcc Ala	cac His	tac Tyr 585	att Ile	gac Asp	ggc Gly	ccc Pro	cac His 590	tgc Cya	gtc Val	aag Lys	acc Thr	tgc Cys 595	1951
ccg Pro	gca Ala	gga Gly	gtc Val	atg Met 600	gga Gly	gaa Glu	aac Asn	aac Asn	acc Thr 605	ctg Leu	gtc Val	tgg Trp	aag Lys	tac. Tyr 610	gca Ala	1999

gad As <u>r</u>	gco Ala	r Gl <sup>7</sup> s aad	cat His	: Val	j tgo . Cys	cac His	cto Lev	tgo Cys 620	His	cca Pro	a aac o Asr	tgo Cys	aco Thi	с Туз	gga Gly	2047
tgo Cys	act Thr	630 Gl7	Pro	ggt Gly	ctt Leu	gaa Glu	ggc Gly 635	Cys	cca Pro	aco Thi	g aat c Asn	999 Gly 640	Pro	aag Lys	g atc s Ile	2095
ccg Pro	Ser 645	. TTE	gco Ala	act Thr	. GTĀ	atg Met 650	Val	Gly 999	gcc Ala	cto Leu	c ctc Leu 655	Leu	cto Lev	g ct <u>c</u> . Leu	gtg Val	2143
gtg Val 660	Ата	cto Lev	. Gly agg	ato Ile	ggc Gly 665	Leu	ttc Phe	atg Met	cga Arg	agg Arg 670	, Arg	cac His	ato	gtt Val	cgg Arg 675	2191
aag Lys	cgc Arg	acg Thr	teu Leu	cgg Arg 680	Arg	ctg Leu	ctg Leu	cag Gln	gag Glu 685	Arg	gag Glu	ctt Leu	gtg Val	gag Glu 690	Pro	2239
ctt Leu	aca Thr	ccc Pro	agt Ser	gga Gly	gaa Glu	gct Ala	ccc Pro	aac Asn	caa Gln	gct Ala	ctc Leu	ttg Leu	agg Arg	atc Ile	ttg Leu	2287
			695					700					705	•		
aag Lys	gaa Glu	act Thr 710	gaa Glu	ttc Phe	aaa Lys	aag Lys	atc Ile 715	aaa Lys	gtg Val	ctg Leu	ggc	tcc Ser 720	ggt Gly	gcg Ala	ttc Phe	2335
ggc	acg Thr 725	gtg Val	tat Tyr	aag Lys	gga Gly	ctc Leu 730	tgg Trp	atc Ile	cca Pro	gaa Glu	ggt Gly 735	gag Glu	aaa Lys	gtt Val	aaa Lys	2383
att Ile 740	ccc Pro	gtc Val	gct Ala	atc Ile	aag Lys 745	gaa Glu	tta Leu	aga Arg	gaa Glu	gca Ala 750	aca Thr	tct Ser	ccg Pro	aaa Lys	gcc Ala 755	2431
aac Asn	aag Lys	gaa Glu	atc Ile	ctc Leu 760	gat Asp	gaa Glu	gcc Ala	tac Tyr	gtg Val 765	atg Met	gcc Ala	agc Ser	gtg Val	gac Asp 770	aac Asn	2479
ccc Pro	cac His	gtg Val	tgc Cys 775	cgc Arg	ctg Leu	ctg Leu	ggc Gly	atc Ile 780	tgc Cys	ctc Leu	acc Thr	tcc Ser	acc Thr 785	gtg Val	cag Gln	2527
ctc Leu	atc Ile	acg Thr 790	cag Gln	ctc Leu	atg Met	ccc Pro	ttc Phe 795	ggc Gly	tgc Cys	ctc Leu	ctg Leu	gac Asp 800	tat Tyr	gtc Val	cgg Arg	2575
gaa Glu	cac His 805	aaa Lys	gac Asp	aat Asn	att Ile	ggc Gly 810	tcc Ser	cag Gln	tac Tyr	ctg Leu	ctc Leu 815	aac Asn	tgg Trp	tgt Cys	gtg Val	2623
cag Gln 820	atc Ile	gca Ala	aag Lys	ggc Gly	atg Met 825	aac Asn	tac Tyr	ttg Leu	Glu	gac Asp 830	cgt Arg	cgc Arg	ttg Leu	Val	cac His 835	2671
cgc Arg	gac Asp	ctg Leu	MIG	gcc Ala 840	agg Arg	aac Asn	gta Val	Leu	gtg Val 845	aaa Lys	aca Thr	ccg Pro	cag Gln	cat <sup>.</sup> His 850	gtc Val	2719

														•		
														gag Glu		2767
														gca Ala		2815
														tgg Trp		2863
														cca Pro		2911
														gga Gly 930		2959
														atc Ile		3007
														cgt Arg		3055
ttg Leu	atc Ile 965	atc Ile	gaa Glu	ttc Phe	tcc Ser	aaa Lys 970	atg Met	gcc Ala	cga Arg	gac Asp	ccc Pro 975	cag Gln	cgc Arg	tac Tyr	ctt Leu	3103
gtc Val 980	att Ile	cag Gln	Gly 333	gat Asp	gaa Glu 985	aga Arg	atg Met	cat His	ttg Leu	cca Pro 990	agt Ser	cct Pro	aca Thr	gac Asp	tcc Ser 995	3151
aac Asn	ttc Phe	tac Tyr	Arg	gcc Ala 1000	ctg Leu	atg Met	gat Asp	Glu	gaa Glu .005	gac Asp	atg Met	gac Asp	Asp	gtg Val 1010	gtg Val	3199
gat Asp	gcc Ala	Asp	gag Glu .015	tac Tyr	ctc Leu	atc Ile	Pro	cag Gln 1020	cag Gln	Gly	ttc Phe	Phe	agc Ser 1025	agc Ser	ccc Pro	3247
	Thr					Leu					Ser			agc Ser		3295
Asn					Cys					Gly				tgt Cys		3343
	Lys			Ser					Tyr					aca Thr 1		3391
gcc Ala	ttg Leu	act Thr	Glu	gac Asp .080	agc Ser	ata Ile	gac Asp	Asp	acc Thr .085	ttc Phe	ctc Leu	cca Pro	Val	cct Pro .090	gaa Glu	3439
tac	ata	aac	cag	tcc	gtt	ccc	aaa	agg	ccc	gct	ggc	tct	gtg	cag	aat	3487

Tyr Ile Asn Gln Ser Val Pro Lys Arg Pro Ala Gly Ser Val Gln Asn 1095 1100 1105	
cct gtc tat cac aat cag cct ctg aac ccc gcg ccc agc aga gac cca Pro Val Tyr His Asn Gln Pro Leu Asn Pro Ala Pro Ser Arg Asp Pro 1110 1115 1120	3535
cac tac cag gac ccc cac agc act gca gtg ggc aac ccc gag tat ctc His Tyr Gln Asp Pro His Ser Thr Ala Val Gly Asn Pro Glu Tyr Leu 1125 1130 1135	3583
aac act gtc cag ccc acc tgt gtc aac agc aca ttc gac agc cct gcc Asn Thr Val Gln Pro Thr Cys Val Asn Ser Thr Phe Asp Ser Pro Ala 1140 1145 1150 1155	8631
cac tgg gcc cag aaa ggc agc cac caa att agc ctg gac aac cct gac. 3 His Trp Ala Gln Lys Gly Ser His Gln Ile Ser Leu Asp Asn Pro Asp 1160 1165 1170	679
tac cag cag gac ttc ttt ccc aag gaa gcc aag cca aat ggc atc ttt 3 Tyr Gln Gln Asp Phe Phe Pro Lys Glu Ala Lys Pro Asn Gly Ile Phe 1175 1180 1185	727
aag ggc tcc aca gct gaa aat gca gaa tac cta agg gtc gcg cca caa 3 Lys Gly Ser Thr Ala Glu Asn Ala Glu Tyr Leu Arg Val Ala Pro Gln 1190 1195 1200	775
agc agt gaa ttt att gga gca tga ccacggagga tagtatgagc cctaaaaatc 3 Ser Ser Glu Phe Ile Gly Ala 1205 1210	829
cagactettt egatacecag gaccaageca cageaggtee tecateccaa cagecatgee 3	889
cgcattagct cttagaccca cagactggtt ttgcaacgtt tacaccgact agccaggaag 3	949
tacttccacc tcgggcacat tttgggaagt tgcattcctt tgtcttcaaa ctgtgaagca 4	009
tttacagaaa cgcatccagc aagaatattg tccctttgag cagaaattta tctttcaaag 4	069
aggtatattt gaaaaaaaa aaaaaagtat atgtgaggat ttttattgat tggggatctt 4:	129
ggagtttttc attgtcgcta ttgattttta cttcaatggg ctcttccaac aaggaagaag 4:	189
cttgctggta gcacttgcta ccctgagttc atccaggccc aactgtgagc aaggagcaca 42	249
agccacaagt cttccagagg atgcttgatt ccagtggttc tgcttcaagg cttccactgc 43	309
aaaacactaa agatccaaga aggccttcat ggccccagca ggccggatcg gtactgtatc 43	369
aagtcatggc aggtacagta ggataagcca ctctgtccct tcctgggcaa agaagaaacg 44	129
gaggggatga attetteett agaettaett ttgtaaaaat gteeceaegg taettaetee 44	189
ccactgatgg accagtggtt tccagtcatg agcgttagac tgacttgttt gtcttccatt 45	
	549
ccattgtttt gaaactcagt atgccgcccc tgtcttgctg tcatgaaatc agcaagagag 46	
	509

agaagactac aaaaatgaag ctgctctgaa atctccttta gccatcaccc caacccccca 4849
aaattagttt gtgttactta tggaagatag ttttctcctt ttacttcact tcaaaagctt 4909
tttactcaaa gagtatatgt tccctccagg tcagctgccc ccaaaccccc tccttacgct 4969
ttgtcacaca aaaagtgtct ctgccttgag tcatctattc aagcacttac agctctggcc 5029
acaacagggc attttacagg tgcgaatgac agtagcatta tgagtagtgt gaattcaggt 5089
agtaaatatg aaactagggt ttgaaattga taatgctttc acaacatttg cagatgttt 5149
agaaggaaaa aagttccttc ctaaaataat ttctctacaa ttggaagatt ggaagattca 5209
gctagttagg agcccatttt ttcctaatct gtgtgtgccc tgtaacctga ctggttaaca 5269
gcagtccttt gtaaacagtg ttttaaactc tcctagtcaa tatccaccc atccaattta 5329
tcaaggaaga aatggttcag aaaatattt cagcctacag ttatgttcag tcacacaca 5389
atacaaaatg ttccttttgc ttttaaagta attttgact cccagatcag tcagagccc 5449
tacagcattg ttaagaaagt atttgattt tgtctcaatg aaaataaac tatattcatt 5509
tcc

<210> 22

<211> 1210

<212> PRT

<213> Homo sapiens

<223> Human EGFR

<400> 22

Met Arg Pro Ser Gly Thr Ala Gly Ala Ala Leu Leu Ala Leu Leu Ala Ala Leu Cys Pro Ala Ser Arg Ala Leu Glu Glu Lys Lys Val Cys Gln 25 Gly Thr Ser Asn Lys Leu Thr Gln Leu Gly Thr Phe Glu Asp His Phe Leu Ser Leu Gln Arg Met Phe Asn Asn Cys Glu Val Val Leu Gly Asn Leu Glu Ile Thr Tyr Val Gln Arg Asn Tyr Asp Leu Ser Phe Leu Lys Thr Ile Gln Glu Val Ala Gly Tyr Val Leu Ile Ala Leu Asn Thr Val Glu Arg Ile Pro Leu Glu Asn Leu Gln Ile Ile Arg Gly Asn Met Tyr Tyr Glu Asn Ser Tyr Ala Leu Ala Val Leu Ser Asn Tyr Asp Ala Asn 120 125 Lys Thr Gly Leu Lys Glu Leu Pro Met Arg Asn Leu Gln Glu Ile Leu 135 140 His Gly Ala Val Arg Phe Ser Asn Asn Pro Ala Leu Cys Asn Val Glu 150 155 Ser Ile Gln Trp Arg Asp Ile Val Ser Ser Asp Phe Leu Ser Asn Met 170 Ser Met Asp Phe Gln Asn His Leu Gly Ser Cys Gln Lys Cys Asp Pro

Ser Cys Pro Asn Gly Ser Cys Trp Gly Ala Gly Glu Glu Asn Cys Gln Lys Leu Thr Lys Ile Ile Cys Ala Gln Gln Cys Ser Gly Arg Cys Arg Gly Lys Ser Pro Ser Asp Cys Cys His Asn Gln Cys Ala Ala Gly Cys Thr Gly Pro Arg Glu Ser Asp Cys Leu Val Cys Arg Lys Phe Arg Asp Glu Ala Thr Cys Lys Asp Thr Cys Pro Pro Leu Met Leu Tyr Asn Pro Thr Thr Tyr Gln Met Asp Val Asn Pro Glu Gly Lys Tyr Ser Phe Gly Ala Thr Cys Val Lys Lys Cys Pro Arg Asn Tyr Val Val Thr Asp His Gly Ser Cys Val Arg Ala Cys Gly Ala Asp Ser Tyr Glu Met Glu Glu Asp Gly Val Arg Lys Cys Lys Cys Glu Gly Pro Cys Arg Lys Val Cys Asn Gly Ile Gly Ile Gly Glu Phe Lys Asp Ser Leu Ser Ile Asn Ala Thr Asn Ile Lys His Phe Lys Asn Cys Thr Ser Ile Ser Gly Asp Leu His Ile Leu Pro Val Ala Phe Arg Gly Asp Ser Phe Thr His Thr Pro Pro Leu Asp Pro Gln Glu Leu Asp Ile Leu Lys Thr Val Lys Glu Ile Thr Gly Phe Leu Leu Ile Gln Ala Trp Pro Glu Asn Arg Thr Asp Leu His Ala Phe Glu Asn Leu Glu Ile Ile Arg Gly Arg Thr Lys Gln His Gly Gln Phe Ser Leu Ala Val Val Ser Leu Asn Ile Thr Ser Leu Gly Leu Arg Ser Leu Lys Glu Ile Ser Asp Gly Asp Val Ile Ile Ser Gly Asn Lys Asn Leu Cys Tyr Ala Asn Thr Ile Asn Trp Lys Lys Leu Phe Gly Thr Ser Gly Gln Lys Thr Lys Ile Ile Ser Asn Arg Gly Glu Asn Ser Cys Lys Ala Thr Gly Gln Val Cys His Ala Leu Cys Ser Pro Glu Gly Cys Trp Gly Pro Glu Pro Arg Asp Cys Val Ser Cys Arg Asn Val Ser Arg Gly Arg Glu Cys Val Asp Lys Cys Asn Leu Leu Glu Gly Glu Pro Arg Glu Phe Val Glu Asn Ser Glu Cys Ile Gln Cys His Pro Glu Cys Leu Pro Gln Ala Met Asn Ile Thr Cys Thr Gly Arg Gly Pro Asp Asn Cys Ile Gln Cys Ala His Tyr Ile Asp Gly Pro His Cys Val Lys Thr Cys Pro Ala Gly Val Met Gly Glu Asn Asn Thr Leu Val Trp Lys Tyr Ala Asp Ala Gly His Val Cys His Leu Cys His Pro Asn Cys Thr Tyr Gly Cys Thr Gly Pro Gly Leu Glu Gly Cys Pro Thr Asn Gly Pro Lys Ile Pro Ser Ile Ala Thr Gly Met Val Gly Ala Leu Leu Leu Leu Val Val Ala Leu Gly Ile Gly Leu Phe Met Arg Arg Arg His Ile Val Arg Lys Arg Thr Leu Arg Arg Leu Leu Gln Glu Arg Glu Leu

	•	675					680					685			
Val	Glu 690	Pro	Leu	Thr	Pro	Ser 695	Gly		Ala	Pro	Asn 700			Leu	Leu
Arg 705	Ile		Lys	Glu	Thr 710		Phe	Lys	Lys	Ile 715		Val	Leu	Gly	
		Phe	Gly	Thr 725		Tyr	Lys	Gly	Leu 730	Trp	Ile	Pro	Glu	Gly 735	720 Glu
Lys	Val	Lys	Ile 740	Pro	Val	Ala	Ile	Lys 745			Arg	Glu	Ala 750		Ser
Pro	Lys	Ala 755			Glu	Ile	Leu 760		Glu	Ala	Tyr	Val 765	Met	Ala	Ser
Val	Asp 770	Asn	Pro	His	Val	Cys 775	Arg	Leu	Leu	Gly	Ile 780		Leu	Thr	Ser
Thr 785	Val	Gln	Leu	Ile	Thr 790	Gln	Leu	Met	Pro	Phe 795		Cys	Leu	Leu	Asp 800
				805			Asn		810	Ser				815	Asn
			820				Gly	825					830		_
		835					Ala 840		-			845			
	850					855	Phe				860				
865					870		Glu			875					880
				885			His		890					895	
			900				Val	905					910	_	
		915					Ala 920					925			
	930					935	Pro				940	•			-
945					950		Met			955			_		960
				965			Phe		970					975	
			980				Asp	985					990		
		995				1	Ala 000				1	.005			
1	.010				1	.015	Tyr			1	.020				
1025	er.	PLO	261	1117	.030	Arg	Thr	Pro		Leu 1035	Ser	Ser	Leu		
		Asn	Asn 1			Val	Ala				Arg	Asn		Leu .055	.040 Gln
Ser	Cys	Pro 1			Glu	Asp	Ser 1			Gln	Arg		Ser .070	Ser	Asp
	1	.075				1	Asp 080	Ser			1	Thr 085	Phe		
1	.090				1	Gln 095	Ser			1	Arg	Pro			
1105				1	.110		Asn		1	Leu .115	Asn			1	120
Arg			1	125				1	Ser 130	Thr			1	Asn 135	Pro
Glu		1	140				1	145				1	Thr 150	Phe-	_
Ser	Pro 1	Ala 155	His	Trp	Ala	Gln 1	Lys 160	Gly	Ser	His		Ile 165	Ser	Leu	Asp

Asn Pro Asp Tyr Gln Gln Asp Phe Phe Pro Lys Glu Ala Lys Pro Asn 1170 1175 1180

Gly Ile Phe Lys Gly Ser Thr Ala Glu Asn Ala Glu Tyr Leu Arg Val 1185 1190 1195 1200

Ala Pro Gln Ser Ser Glu Phe Ile Gly Ala 1210

<210> 23 <211>. 5935 <212> DNA <213> Mus musculus <220> <221> CDS <222> (224)..(3856) <220> <223> Murine EGFR <400> 23 gaatteggge ceteetette treeegeact gtgegeteet eetgggetag ggegtetgga 60 tegagteceg gagetacege eteccagaca gaegaegggt cacetggaeg egageetgtg 120 teegggtete gtegttgeeg gegeagteae tgggeacaae egtgggaete egtetgtete 180 ggattaatcc cggagagcca gagccaacct ctcccggtca gag atg cga ccc tca Met Arg Pro Ser ggg acc gcg aga acc aca ctg ctg gtg ctg ctg acc gcg ctc tgc gcc 283 Gly Thr Ala Arg Thr Thr Leu Leu Val Leu Leu Thr Ala Leu Cys Ala 10 gca ggt ggg gcg ttg gag gaa aag aaa gtc tgc caa ggc aca agt aac 331 Ala Gly Gly Ala Leu Glu Glu Lys Lys Val Cys Gln Gly Thr Ser Asn 25 agg ctc acc caa ctg ggc act ttt gaa gac cac ttt ctg agc ctg cag 379 Arg Leu Thr Gln Leu Gly Thr Phe Glu Asp His Phe Leu Ser Leu Gln 40 45 agg atg tac aac aac tgt gaa gtg gtc ctt ggg aac ttg gaa att acc Arg Met Tyr Asn Asn Cys Glu Val Val Leu Gly Asn Leu Glu Ile Thr 55 60 tat gtg caa agg aat tac gac ctt tcc ttc tta aag acc atc cag gag Tyr Val Gln Arg Asn Tyr Asp Leu Ser Phe Leu Lys Thr Ile Gln Glu 70 gtg gcc ggc tat gtc ctc att gcc ctc aac acc gtg gag aga atc cct 523 Val Ala Gly Tyr Val Leu Ile Ala Leu Asn Thr Val Glu Arg Ile Pro 85 90 ttg gag aac ctg cag atc atc agg gga aat gct ctt tat gaa aac acc 571 Leu Glu Asn Leu Gln Ile Ile Arg Gly Asn Ala Leu Tyr Glu Asn Thr 105 115 tat gcc tta gcc atc ctg tcc aac tat ggg aca aac aga act ggg ctt

Tyr	Ala	Leu	Ala 120	Ile	Leu	Ser	Asn	Tyr 125	Gly	Thr	Asn	Arg	Thr 130		Leu	
agg Arg	gaa Glu	ctg Leu 135	ccc	atg Met	cgg Arg	aac Asn	tta Leu 140	cag Gln	gaa Glu	atc Ile	ctg Leu	att Ile 145	ggt Gly	gct Ala	gtg Val	667
cga Arg	ttc Phe 150	agc Ser	aac Asn	aac Asn	ccc Pro	atc Ile 155	ctc Leu	tgc Cys	aat Asn	atg Met	gat Asp 160	act Thr	atc Ile	cag Gln	tgg Trp	715
agg Arg 165	gac Asp	atc Ile	gtc Val	caa Gln	aac Asn 170	gtc Val	ttt Phe	atg Met	agc Ser	aac Asn 175	atg Met	tca Ser	atg Met	gac Asp	tta Leu 180	763
cag Gln	agc Ser	cat His	ccg Pro	agc Ser 185	agt Ser	tgc Cys	ccc Pro	aaa Lys	tgt Cys 190	gat Asp	cca Pro	agc Ser	tgt Cys	ccc Pro 195	aat Asn	811
gga Gly	agc Ser	tgc Cys	tgg Trp 200	gga Gly	gga Gly	gga Gly	gag Glu	gag Glu 205	aac Asn	tgc Cys	cag Gln	aaa Lys	ttg Leu 210	acc Thr	aaa Lys	859
atc Ile	atc Ile	tgt Cys 215	gcc Ala	cag Gln	caa Gln	tgt Cys	tcc Ser 220	cat His	cgc Arg	tgt Cys	cgt Arg	ggc Gly 225	agg Arg	tcc Ser	ccc Pro	907
agt Ser	gac Asp 230	tgc Cys	tgc Cys	cac His	aac Asn	caa Gln 235	tgt Cys	gct Ala	gcg Ala	gl <sup>y</sup> aaa	tgt Cys 240	aca Thr	Gly aaa	ccc Pro	cga Arg	955
gag Glu 245	agt Ser	gac Asp	tgt Cys	ctg Leu	gtc Val 250	tgc Cys	caa Gln	aag Lys	ttc Phe	caa Gln 255	gat Asp	gag Glu	gcc Ala	aca Thr	tgc Cys 260	1003
aaa Lys	gac Asp	acc Thr	tgc Cys	cca Pro 265	cca Pro	ctc Leu	atg Met	ctg Leu	tac Tyr 270	aac Asn	ccc Pro	acc Thr	acc Thr	tat Tyr 275	cag Gln	1051
atg Met	gat Asp	gtc Val	aac Asn 280	cct Pro	gaa Glu	gjå aaa	aag Lys	tac Tyr 285	agc Ser	ttt. Phe	ggt Gly	gcc Ala	acc Thr 290	tgt Cys	gtg Val	1099
aag Lys	aag Lys	tgc Cys 295	ccc Pro	cga Arg	aac Asn	tac Tyr	gtg Val 300	gtg Val	aca Thr	gat Asp	cat His	ggc Gly 305	tca Ser	tgt Cys	gtc Val	1147
cga Arg	gcc Ala 310	tgt Cys	Gly 999	cct Pro	gac Asp	tac Tyr 315	tac Tyr	gaa Glu	gtg Val	gaa Glu	gaa Glu 320	gat Asp	ggc Gly	atc Ile	cgc Arg	1195
aag Lys 325	tgt Cys	aaa Lys	aaa Lys	tgt Cys	gat Asp 330	gly aaa	ccc Pro	tgt Cys	cgc Arg	aaa Lys 335	gtt Val	tgt Cys	aat Asn	ggc	ata Ile 340	1243
ggc Gly	att Ile	ggt Gly	gaa Glu	ttt Phe 345	aaa Lys	gac Asp	aca Thr	Leu	tcc Ser 350	ata Ile	aat Asn	gct Ala	aca Thr	aac Asn 355	atc Ile	1291
aaa Lys	cac His	ttc Phe	aaa Lys	tac Tyr	tgc Cys	act Thr	gcc Ala	atc Ile	agc Ser	Gly aaa	gac Asp	ctt Leu	cac His	atc Ile	ctg Leu	1339

1

360 365 370

cca Pro	gtg Val	gcc Ala 375	Phe	aag Lys	gly aaa	gat Asp	tct Ser 380	Phe	acg Thr	cgc Arg	act Thr	cct Pro 385	cct Pro	cta Leu	gac Asp	1387
cca Pro	cga Arg 390	gaa Glu	cta Leu	gaa Glu	att Ile	cta Leu 395	aaa Lys	acc Thr	gta Val	aag Lys	gaa Glu 400	ata Ile	aca Thr	ggc	ttt Phe	1435
ttg Leu 405	ctg Leu	att Ile	cag Gln	gct Ala	tgg Trp 410	cct Pro	gat Asp	aac Asn	tgg Trp	act Thr 415	gac Asp	ctc Leu	cat His	gct Ala	ttc Phe 420	1483
gag Glu	aac Asn	cta Leu	gaa Glu	ata Ile 425	ata Ile	cgt Arg	Gly	aga Arg	aca Thr 430	aag Lys	caa Gln	cat His	ggt Gly	cag Gln 435	ttt Phe	1531
tct Ser	ttg Leu	gcg Ala	gtc Val 440	gtt Val	ggc	ctg Leu	aac Asn	atc Ile 445	Thr	tca Ser	ctg Leu	Gly 999	ctg Leu 450	cgt Arg	tcc Ser	1579
ctc Leu	aag Lys	gag Glu 455	atc Ile	agt Ser	gat Asp	Gly	gat Asp 460	gtg Val	atc Ile	att Ile	tct Ser	gga Gly 465	aac Asn	cga Arg	aat Asn	1627
ttg Leu	tgc Cys 470	tac Tyr	gca Ala	aac Asn	aca Thr	ata Ile 475	aac Asn	tgg Trp	aaa Lys	aaa Lys	ctc Leu 480	ttc Phe	ely aaa	aca Thr	ccc Pro	1675
aat Asn 485	cag Gln	aaa Lys	acc Thr	aaa Lys	atc Ile 490	atg Met	aac Asn	aac Asn	aga Arg	gct Ala 495	gag Glu	aaa Lys	gac Asp	tgc Cys	aag Lys 500	1723
gcc Ala	gtg Val	aac Asn	cac His	gtc Val 505	tgc Cys	aat Asn	cct Pro	tta Leu	tgc Cys 510	tcc Ser	tcg Ser	gaa Glu	ggc	tgc Cys 515	tgg Trp	1771
ggc Gly	cct Pro	gag Glu	ccc Pro 520	agg Arg	gac Asp	tgt Cys	gtc Val	tcc Ser 525	tgc Cys	cag Gln	aat Asn	gtg Val	agc Ser 530	aga Arg	ggc	1819
agg Arg	gag Glu	tgc Cys 535	gtg Val	gag Glu	aaa Lys	tgc Cys	aac Asn 540	atc Ile	ctg Leu	gag Glu	Gly 333	gaa Glu 545	cca Pro	agg Arg	gag Glu	1867
ttt Phe	gtg Val 550	gaa Glu	aat Asn	tct Ser	gaa Glu	tgc Cys 555	atc Ile	cag Gln	tgc Cys	cat His	cca Pro 560	gaa Glu	tgt Cys	ctg Leu	ccc Pro	1915
cag Gln 565	gcc Ala	atg Met	aac Asn	atc Ile	acc Thr 570	tgt Cys	aca Thr	ggc Gly	agg Arg	999 Gly 575	cca Pro	gac Asp	aac Asn	tgc Cys	atc Ile 580	1963
cag Gln	tgt Cys	gcc Ala	cac His	tac Tyr 585	att Ile	gat Asp	ggc	cca Pro	cac His 590	tgt Cys	gtc Val	aag Lys	acc Thr	tgc Cys 595	cca Pro	2011
gct Ala	ggc	Ile	atg Met 600	gga Gly	gag Glu	aac Asn	Asn	act Thr 605	ctg Leu	gtc Val	tgg Trp	aag Lys	tat Tyr 610	gca Ala	gat Asp	2059

gcc Ala	aat Asn	aat Asn 615	gtc Val	tgc Cys	cac His	cta Leu	tgc Cys 620	cac His	gcc Ala	aac Asn	tgt Cys	acc Thr 625	tat Tyr	gga Gly	tgt Cys	2107
gct Ala	630 Gly 999	cca Pro	ggt Gly	ctt Leu	caa Gln	gga Gly 635	tgt Cys	gaa Glu	gtg Val	tgg Trp	cca Pro 640	tct Ser	Gly 999	cca Pro	aag Lys	2155
ata Ile 645	cca Pro	tct Ser	att Ile	gcc Ala	act Thr 650	GJÀ 888	att Ile	gtg Val	ggt Gly	ggc Gly 655	ctc Leu	ctc Leu	ttc Phe	ata Ile	gtg Val 660	2203
gtg Val	gtg Val	gcc Ala	ctt Leu	665 Gly 333	att Ile	ggc	cta Leu	ttc Phe	atg Met 670	cga Arg	aga Arg	cgt Arg	cac His	att Ile 675	gtt Val	2251
cga Arg	aag Lys	cgt Arg	aca Thr 680	cta Leu	cgc Arg	cgc Arg	ctg Leu	ctt Leu 685	caa Gln	gag Glu	aga Arg	Glu	ctc Leu .690	gtg Val	gaa Glu	2299
cct Pro	ctc Leu	aca Thr 695	ccc Pro	agc Ser	gga Gly	gaa Glu	gct Ala 700	cca Pro	aac Asn	caa Gln	gcc Ala	cac His 705	ttg Leu	agg Arg	ata Ile	2347
tta Leu	aag Lys 710	gaa Glu	aca Thr	gaa Glu	ttc Phe	aaa Lys 715	aag Lys	atc Ile	aaa Lys	gtt Val	ctg Leu 720	ggt Gly	tcg Ser	gga Gly	gca Ala	2395
ttt Phe 725	ggc Gly	aca Thr	gtg Val	tat Tyr	aag Lys 730	ggt Gly	ctc Leu	tgg Trp	atc Ile	cca Pro 735	gaa Glu	ggt Gly	gag Glu	aaa Lys	gta Val 740	2443
aaa Lys	atc Ile	ccg Pro	gtg Val	gcc Ala 745	atc Ile	aag Lys	gag Glu	tta Leu	aga Arg 750	gaa Glu	gcc Ala	aca Thr	·tct Ser	cca Pro 755	aaa Lys	2491
gcc Ala	aac Asn	aaa Lys	gaa Glu 760	atc Ile	ctt Leu	gac Asp	gaa Glu	gcc Ala 765	tat Tyr	gtg Val	atg Met	gct Ala	agt Ser 770	gtg Val	gac Asp	2539
aac Asn	cct Pro	cat His 775	gta Val	tgc Cys	cgc Arg	ctc Leu	ctg Leu 780	ggc	atc Ile	tgt Cys	ctg Leu	acc Thr 785	tcc Ser	act Thr	gtc Val	2587
cag Gln	ctc Leu 790	att Ile	aca Thr	cag Gln	ctc Leu	atg Met 795	ccc Pro	tac Tyr	ggt Gly	tgc Cys	ctc Leu 800	ctg Leu	gac Asp	tac Tyr	gtc Val	2635
cga Arg 805	gaa Glu	cac His	aag Lys	gac Asp	aac Asn 810	att Ile	Gly	tcc Ser	cag Gln	tac Tyr 815	ctc Leu	ctc Leu	aac Asn	tgg Trp	tgt Cys 820	2683
gtg Val	cag Gln	att Ile	gca Ala	aag Lys 825	ggc	atg Met	aac Asn	tac Tyr	ctg Leu 830	gaa Glu	gat Asp	cgg Arg	cgt Arg	ttg Leu 835	gtg Val	2731
cac His	cgt Arg	gac Asp	ttg Leu 840	gca Ala	gcc Ala	agg Arg	aat Asn	gta Val 845	ctg Leu	gtg Val	aag Lys	aca Thr	cca Pro 850	cag Gln	cat His	2779

gtc Val	aag Lys	atc Ile 855	aca Thr	gat Asp	ttt Phe	ggg Gly	ctg Leu 860	gcc Ala	aaa Lys	ctg Leu	ctt Leu	ggt Gly 865	gct Ala	gaa Glu	gag Glu	2827
aaa Lys	gaa Glu	tat Tyr	cat His	gcc Ala	gag Glu	Gly 999	ggc	aaa Lys	gtg Val	cct Pro	atc Ile	aag Lys	tgg Trp	atg Met	gct Ala	28 <b>7</b> 5
٠	870					875					880					
ttg Leu 885	gaa Glu	tca Ser	att Ile	tta Leu	cac His 890	cga Arg	att Ile	tat Tyr	aca Thr	cac His 895	caa Gln	agt Ser	gat Asp	gtc Val	tgg Trp 900	2923
agc Ser	tat Tyr	ggt Gly	gtc Val	act Thr 905	gtg Val	tgg Trp	gaa Glu	ctg Leu	atg Met 910	acc Thr	ttt Phe	GJA aaa	tcc Ser	aag Lys 915	cct Pro	2971
tat Tyr	gat Asp	gga Gly	atc Ile 920	cca Pro	gca Ala	agt Ser	gac Asp	atc Ile 925	tca Ser	tcc Ser	atc Ile	Leu	gag Glu 930	aaa Lys	gga Gly	3019
gaa Glu	cgc Arg	ctt Leu 935	cca Pro	cag Gln	cca Pro	cct Pro	atc Ile 940	tgc Cys	acc Thr	atc Ile	gat Asp	gtc Val 945	tac Tyr	atg Met	atc Ile	3067
atg Met	gtc Val 950	aag Lys	tgc Cys	tgg Trp	atg Met	ata Ile 955	gat Asp	gct Ala	gat Asp	agc Ser	cgc Arg 960	cca Pro	aag Lys	ttc Phe	cga Arg	3115
gag Glu 965	ttg Leu	att Ile	ctt Leu	gaa Glu	ttc Phe 970	tcc Ser	aaa Lys	atg Met	gcc Ala	cga Arg 975	gac Asp	cca Pro	cag Gln	cgc Arg	tac Tyr 980	3163
ctt Leu	gtt Val	atc Ile	cag Gln	999 Gly 985	gat Asp	gaa Glu	aga Arg	atg Met	cat His 990	ttg Leu	cca Pro	agc Ser	cct Pro	aca Thr 995	gac Asp	3211
tcc Ser	aac Asn	Phe	tac Tyr 1000	cga Arg	gcc Ala	ctg Leu	Met	gat Asp .005	gaa Glu	gag Glu	gac Asp	Met	gag Glu 1010	gat Asp	gta Val	3259
gtt Val	Asp	gct Ala 1015	gat Asp	gag Glu	tat Tyr	Leu	acc Thr 1020	cca Pro	cag Gln	caa Gln	ggc ggc	ttc Phe .025	ttc Phe	aac Asn	agc Ser	3307
Pro	tcc Ser .030	acg Thr	tcg Ser	agg Arg	Thr	ccc Pro .035	ctc Leu	ttg Leu	agt Ser	Ser	ctg Leu .040	agt Ser	gca Ala	act Thr	agc Ser	3355
aac Asn 1045	Asn	tcc Ser	act Thr	Val	gct Ala .050	tgc Cys	att Ile	aat Asn	Arg	aat Asn .055	gjå aaa	agc Ser	tgc Cys	Arg	gtc Val .060	3403
aaa Lys	gaa Glu	gac Asp	Ala	ttc Phe 065	ttg Leu	cag Gln	cgg Arg	Tyr	agc Ser 070	tcc Ser	gac Asp	ccc Pro	Thr	ggt Gly .075	gct Ala	3451
gta Val	aca Thr	Glu	gac Asp .080	aac Asn	ata Ile	gat Asp	Asp	gca Ala 085	ttc Phe	ctc Leu	cct Pro	Val	cct Pro 090	gaa Glu	tat Tyr	3499

gta aac caa tct gtt c Val Asn Gln Ser Val P 1095	cc aag agg ro Lys Arg 1100	Pro Ala G	gc tct gtg ( ly Ser Val ( 1105	Cag aac cct Gln Asn Pro	3547
gtc tat cac aat cag c Val Tyr His Asn Gln P 1110	cc ctg cat ro Leu His 1115	cca gct co Pro Ala Pr	et gga aga g o Gly Arg 1 1120	gac ctg cat Asp Leu His	3595
tat caa aat ccc cac a Tyr Gln Asn Pro His So 1125 11:	er Asn Ala	gtg ggc aa Val Gly As 113	n Pro Glu I	at ctc aac Tyr Leu Asn 1140	3643
act gcc cag cct acc to Thr Ala Gln Pro Thr Cy 1145	gt ctc agt ys Leu Ser	agt ggg tt Ser Gly Ph 1150	t aac agc o e Asn Ser E	ct gca ctc Pro Ala Leu 1155	3691
tgg atc cag aaa ggc ag Trp Ile Gln Lys Gly Se 1160	er His Gln	atg agc ct Met Ser Le L165	u Asp Asn P	ct gac tac ro Asp Tyr 70	3739
cag cag gac ttc ttc co Gln Gln Asp Phe Phe Pr 1175	cc aag gaa co Lys Glu 1180	acc aag cc Thr Lys Pr	a aat ggc a o Asn Gly I 1185	ta ttt aag le Phe Lys	3787
ggc ccc aca gct gaa aa Gly Pro Thr Ala Glu As 1190	at gca gag sn Ala Glu 1195	tac cta cgg Tyr Leu Arg	g gtg gca c g Val Ala P 1200	ct cca agc ro Pro Ser	3835
agc gag ttt att gga gc Ser Glu Phe Ile Gly Al 1205 121	a	gaagggg cato	catacca gct	ataaaat	3886
gtctggactt tctagaatcc	caggaccaac	tatggcagca	a cctccactt	c tggtagccat	3946
gcccacgctg tgtcaaatgt	cactcagact	ggctttaaag	g cataactct	g atgggctttg	4006
tcactgagcc aagaagtggg	catatataat	gatgcacttt	gggaagttga	a aggtacatca	4066
attgatette gaactgtgaa	gattccacaa	aaaaggtato	catcgagaac	attgtccatt	4126
ggaacagaag tttgcctcat	ggtgaggtac	atatgggaaa	aaaacagaca	tatggagctt	4186
atgtttaggg aactttggga (	ttettgtett	tattgatttg	attgatgcad	tcttgtagtc	4246
tggtacacag agttgcctgg a					
aagacacttc cgtggcaaga o					
ccacacttgt acagcattaa a					
ctggggctgg gagaagagga a					
aaatatgtcc ctggcaccta a					
ctatgttttc tgtttcattg t					
agtaaacaaa caaacaaaaa a					
cattcgaacc cattcaaacc a					
accatgcaaa aatccataga g					

tgaggtgtgt atatgagact acgaaagtga actactcttc aaatccactt tgccttcact 4846 cctctatacc ctaaatctag tgtaaaccac acatggagga taactttttt ttttaatttt 4906 aaaagtgttt attagatatg tttttcttcc tggtaaactg cagccaaaca tcagttaaga 4966 gccatttttg ataaacacta tcacaatgat ctcgggatcc atcctttccg atttaccaag 5026 tgatggatag acgtgaactc ataaacacta cccataagac aaaacaatga gtgccagaca 5086 agacatcage caggeaccag ageacagage aggactggge aatetgttgg agatatetag 5146 aaagttcaca aaggaaacaa gattgtccac taccttgtga gatctagcag tcataaatac 5206 cagggaaatg gaaagtgtgt ttccttacag caccaggtct tcgatcttcc taatgctgtg 5266 accetttaat acagtttgcc atgttgtggt gacceccaac cataaaatta tttttgttgc 5326 tacttcataa ctgtaaattt gctactctta cagaccacaa tgtaaatatc tgatatgcta 5386 tetgatatge aggetatetg acagaggteg caaccegcag gttgagagee actgeettea 5446 aggetttaat caagagagta gtgagetgag ggetttaetg gtaagteagg ggeaagteea 5506 actcaatcat cctcacatac tggctgctcc ctcaggcctg agaatgaggc ttgcagcatc 5566 ctctggtttc ctaaccgtta tccatccctg actctcatct ctgaaaatag atgtcatcca 5626 tgaaattaag gagtgagaat attaagcagc atttatagag ctcaaaattc catgtcatca 5686 ccaggaagtg ccatgttgat cacagagaac acagaggaga catatagaca gggttttgct 5746 caaaattggg atatagaatg agcctgtcag gtacctatca ggagcggtaa tccgtgagag 5806 agaaccgttg caagccactc taactgtagc aatgaaaccc tagtattttt gtactttgaa 5866 atactttctt ataacaaaat aaagtagcaa aaaaactgtt caaaaaaaaa aaaaaaaaa 5926 cccgaattc 5935

<210> 24 <211> 1210 <212> PRT <213> Mus musculus <223> Murine EGFR

<400> 24

 Met
 Arg
 Pro
 Ser
 Gly
 Thr
 Ala
 Arg
 Thr
 Thr
 Leu
 Leu
 Leu
 Leu
 Leu
 Thr
 Leu
 Leu
 Leu
 Glu
 Leu
 Glu
 Lys
 Lys
 Val
 Cys
 Gln

 Ala
 Leu
 Gly
 Gly
 Ala
 Leu
 Gly
 Thr
 Phe
 Glu
 Asp
 His
 Phe

 Gly
 Thr
 Ser
 Asn
 Arg
 Leu
 Thr
 Asn
 Asn
 Cys
 Glu
 Val
 Leu
 Gly
 Asn

 Leu
 Glu
 Ile
 Thr
 Tyr
 Val
 Asn
 Tyr
 Asp
 Leu
 Leu
 Lys

 65
 70
 75
 75
 80
 80
 Thr
 Val
 Asn
 Thr
 Val
 Leu
 Asn
 Thr
 Val
 Leu
 Asn
 Thr
 Val
 Asn
 Thr
 Thr
 Thr
 Val
 Asn
 Thr

```
Glu Arg Ile Pro Leu Glu Asn Leu Gln Ile Ile Arg Gly Asn Ala Leu
                                 105
 Tyr Glu Asn Thr Tyr Ala Leu Ala Ile Leu Ser Asn Tyr Gly Thr Asn
                             120
 Arg Thr Gly Leu Arg Glu Leu Pro Met Arg Asn Leu Gln Glu Ile Leu
                        135
Ile Gly Ala Val Arg Phe Ser Asn Asn Pro Ile Leu Cys Asn Met Asp
                     150
                                        155
 Thr Ile Gln Trp Arg Asp Ile Val Gln Asn Val Phe Met Ser Asn Met
                 165
                                     170
 Ser Met Asp Leu Gln Ser His Pro Ser Ser Cys Pro Lys Cys Asp Pro
            180
                                 185
Ser Cys Pro Asn Gly Ser Cys Trp Gly Gly Glu Glu Asn Cys Gln
                             200
Lys Leu Thr Lys Ile Ile Cys Ala Gln Gln Cys Ser His Arg Cys Arg
                         215
                                             220
Gly Arg Ser Pro Ser Asp Cys Cys His Asn Gln Cys Ala Ala Gly Cys
                    230
Thr Gly Pro Arg Glu Ser Asp Cys Leu Val Cys Gln Lys Phe Gln Asp
                245
                                     250
Glu Ala Thr Cys Lys Asp Thr Cys Pro Pro Leu Met Leu Tyr Asn Pro
            260
                                 265
Thr Thr Tyr Gln Met Asp Val Asn Pro Glu Gly Lys Tyr Ser Phe Gly
        275
                             280
                                                 285
Ala Thr Cys Val Lys Lys Cys Pro Arg Asn Tyr Val Val Thr Asp His
                        295
                                            300
Gly Ser Cys Val Arg Ala Cys Gly Pro Asp Tyr Tyr Glu Val Glu Glu
                    310
                                        315
Asp Gly Ile Arg Lys Cys Lys Cys Asp Gly Pro Cys Arg Lys Val
                325
                                    330
Cys Asn Gly Ile Gly Ile Gly Glu Phe Lys Asp Thr Leu Ser Ile Asn
                                345
Ala Thr Asn Ile Lys His Phe Lys Tyr Cys Thr Ala Ile Ser Gly Asp
                            360
Leu His Ile Leu Pro Val Ala Phe Lys Gly Asp Ser Phe Thr Arg Thr
                        375
Pro Pro Leu Asp Pro Arg Glu Leu Glu Ile Leu Lys Thr Val Lys Glu
                   390
                                        395
Ile Thr Gly Phe Leu Leu Ile Gln Ala Trp Pro Asp Asn Trp Thr Asp
                405
                                    410
Leu His Ala Phe Glu Asn Leu Glu Ile Ile Arg Gly Arg Thr Lys Gln
            420
                                425
His Gly Gln Phe Ser Leu Ala Val Val Gly Leu Asn Ile Thr Ser Leu
        435
                            440
                                                 445
Gly Leu Arg Ser Leu Lys Glu Ile Ser Asp Gly Asp Val Ile Ile Ser
                        455
                                             460
Gly Asn Arg Asn Leu Cys Tyr Ala Asn Thr Ile Asn Trp Lys Lys Leu
                    470
                                        475
Phe Gly Thr Pro Asn Gln Lys Thr Lys Ile Met Asn Asn Arg Ala Glu
                                    490
Lys Asp Cys Lys Ala Val Asn His Val Cys Asn Pro Leu Cys Ser Ser
                                505
Glu Gly Cys Trp Gly Pro Glu Pro Arg Asp Cys Val Ser Cys Gln Asn
                            520
                                                525
Val Ser Arg Gly Arg Glu Cys Val Glu Lys Cys Asn Ile Leu Glu Gly
                        535
                                            540
Glu Pro Arg Glu Phe Val Glu Asn Ser Glu Cys Ile Gln Cys His Pro
                    550
                                        555
Glu Cys Leu Pro Gln Ala Met Asn Ile Thr Cys Thr Gly Arg Gly Pro
                                    570
Asp Asn Cys Ile Gln Cys Ala His Tyr Ile Asp Gly Pro His Cys Val
```

```
580
                                 585
 Lys Thr Cys Pro Ala Gly Ile Met Gly Glu Asn Asn Thr Leu Val Trp
                            600
                                                605
 Lys Tyr Ala Asp Ala Asn Asn Val Cys His Leu Cys His Ala Asn Cys
                        615
 Thr Tyr Gly Cys Ala Gly Pro Gly Leu Gln Gly Cys Glu Val Trp Pro
                     630
                                        635
 Ser Gly Pro Lys Ile Pro Ser Ile Ala Thr Gly Ile Val Gly Gly Leu
                645
                                     650
 Leu Phe Ile Val Val Val Ala Leu Gly Ile Gly Leu Phe Met Arg Arg
            660
                                665
 Arg His Ile Val Arg Lys Arg Thr Leu Arg Arg Leu Leu Gln Glu Arg
                            680
 Glu Leu Val Glu Pro Leu Thr Pro Ser Gly Glu Ala Pro Asn Gln Ala
                         695
 His Leu Arg Ile Leu Lys Glu Thr Glu Phe Lys Lys Ile Lys Val Leu
                     710
                                         715
 Gly Ser Gly Ala Phe Gly Thr Val Tyr Lys Gly Leu Trp Ile Pro Glu
                 725
                                     730
 Gly Glu Lys Val Lys Ile Pro Val Ala Ile Lys Glu Leu Arg Glu Ala
                                 745
 Thr Ser Pro Lys Ala Asn Lys Glu Ile Leu Asp Glu Ala Tyr Val Met
                             760
 Ala Ser Val Asp Asn Pro His Val Cys Arg Leu Leu Gly Ile Cys Leu
                         775
                                            780
Thr Ser Thr Val Gln Leu Ile Thr Gln Leu Met Pro Tyr Gly Cys Leu
                     790
                                        795
Leu Asp Tyr Val Arg Glu His Lys Asp Asn Ile Gly Ser Gln Tyr Leu
                 805
                                    810
Leu Asn Trp Cys Val Gln Ile Ala Lys Gly Met Asn Tyr Leu Glu Asp
                                825
Arg Arg Leu Val His Arg Asp Leu Ala Ala Arg Asn Val Leu Val Lys
                            840
Thr Pro Gln His Val Lys Ile Thr Asp Phe Gly Leu Ala Lys Leu Leu
                        855
Gly Ala Glu Glu Lys Glu Tyr His Ala Glu Gly Gly Lys Val Pro Ile
                    870
                                        875
Lys Trp Met Ala Leu Glu Ser Ile Leu His Arg Ile Tyr Thr His Gln
                885
                                    890
Ser Asp Val Trp Ser Tyr Gly Val Thr Val Trp Glu Leu Met Thr Phe
            900
                                905
Gly Ser Lys Pro Tyr Asp Gly Ile Pro Ala Ser Asp Ile Ser Ser Ile
                            920
                                                925
Leu Glu Lys Gly Glu Arg Leu Pro Gln Pro Pro Ile Cys Thr Ile Asp
                        935
                                            940
Val Tyr Met Ile Met Val Lys Cys Trp Met Ile Asp Ala Asp Ser Arg
                    950
                                       955
Pro Lys Phe Arg Glu Leu Ile Leu Glu Phe Ser Lys Met Ala Arg Asp
                                   970
Pro Gln Arg Tyr Leu Val Ile Gln Gly Asp Glu Arg Met His Leu Pro
                               985
Ser Pro Thr Asp Ser Asn Phe Tyr Arg Ala Leu Met Asp Glu Glu Asp
                          1000
Met Glu Asp Val Val Asp Ala Asp Glu Tyr Leu Thr Pro Gln Gly
                       1015
                                          1020
Phe Phe Asn Ser Pro Ser Thr Ser Arg Thr Pro Leu Leu Ser Ser Leu
                  1030
                                     1035
Ser Ala Thr Ser Asn Asn Ser Thr Val Ala Cys Ile Asn Arg Asn Gly
              1045 1050
Ser Cys Arg Val Lys Glu Asp Ala Phe Leu Gln Arg Tyr Ser Ser Asp
           1060
                              1065
```

Pro Thr Gly Ala Val Thr Glu Asp Asn Ile Asp Asp Ala Phe Leu Pro 1080 Val Pro Glu Tyr Val Asn Gln Ser Val Pro Lys Arg Pro Ala Gly Ser 1090 1095 1100 Val Gln Asn Pro Val Tyr His Asn Gln Pro Leu His Pro Ala Pro Gly 1110 1115 Arg Asp Leu His Tyr Gln Asn Pro His Ser Asn Ala Val Gly Asn Pro 1125 1130 Glu Tyr Leu Asn Thr Ala Gln Pro Thr Cys Leu Ser Ser Gly Phe Asn 1140 1145 1150 Ser Pro Ala Leu Trp Ile Gln Lys Gly Ser His Gln Met Ser Leu Asp 1160 1165 Asn Pro Asp Tyr Gln Gln Asp Phe Phe Pro Lys Glu Thr Lys Pro Asn 1170 1175 1180 Gly Ile Phe Lys Gly Pro Thr Ala Glu Asn Ala Glu Tyr Leu Arg Val 1190 1195 Ala Pro Pro Ser Ser Glu Phe Ile Gly Ala 1205 1210 <210> 25 <211> 4877 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (443)..(4066) <220> <223> Human EGF <400> 25 actgttggga gaggaatcgt atctccatat ttcttctttc agccccaatc caagggttgt 60 agetggaact ttecateagt tetteettte ttttteetet etaageettt geettgetet 120 gtcacagtga agtcagccag agcagggctg ttaaactctg tgaaatttgt cataagggtg 180 tcaggtattt cttactggct tccaaagaaa catagataaa gaaatctttc ctgtggcttc 240 ccttggcagg ctgcattcag aaggtctctc agttgaagaa agagcttgga ggacaacagc 300 acaacaggag agtaaaagat gccccagggc tgaggcctcc gctcaggcag ccgcatctgg 360 ggtcaatcat actcaccttg cccgggccat gctccagcaa aatcaagctg ttttcttttq 420 aaagttcaaa ctcatcaaga tt atg ctg ctc act ctt atc att ctg ttg cca 472 Met Leu Leu Thr Leu Ile Ile Leu Leu Pro

gta gtt tca aaa ttt agt ttt gtt agt ctc tca gca ccg cag cac tgg 520 Val Val Ser Lys Phe Ser Phe Val Ser Leu Ser Ala Pro Gln His Trp 15 20 25

agc tgt cct gaa ggt act ctc gca gga aat ggg aat tct act tgt gtg 568 Ser Cys Pro Glu Gly Thr Leu Ala Gly Asn Gly Asn Ser Thr Cys Val 30 35 40

ggt Gly	cct Pro	gca Ala 45	Pro	ttc Phe	tta Leu	att Ile	tto Phe 50	Ser	cat His	gga Gly	aat Asn	agt Ser 55	: Ile	ttt Phe	agg Arg	616
att Ile	gac Asp 60	Thr	gaa Glu	gga Gly	acc Thr	aat Asn 65	tat Tyr	gag Glu	caa Gln	ttg Leu	gtg Val 70	gtg Val	gat Asp	gct Ala	ggt Gly	664
gtc Val 75	Ser	gtg Val	atc Ile	atg Met	gat Asp 80	Phe	cat His	tat Tyr	aat Asn	gag Glu 85	Lys	aga Arg	atc Ile	tat Tyr	tgg Trp 90	712
gtg Val	gat Asp	tta Leu	gaa Glu	aga Arg 95	caa Gln	ctt Leu	ttg Leu	caa Gln	aga Arg 100	gtt Val	ttt Phe	ctg Leu	aat Asn	999 Gly 105		760
agg Arg	caa Gln	gag Glu	aga Arg 110	gta Val	tgt Cys	aat Asn	ata Ile	gag Glu 115	aaa Lys	aat Asn	gtt Val	tct Ser	gga Gly 120	atg Met	gca Ala	808
ata Ile	aat Asn	tgg Trp 125	ata Ile	aat Asn	gaa Glu	gaa Glu	gtt Val 130	att Ile	tgg Trp	tca Ser	aat Asn	caa Gln 135	cag Gln	gaa Glu	gga Gly	856
atc Ile	att Ile 140	aca Thr	gta Val	aca Thr	gat Asp	atg Met 145	aaa Lys	gga Gly	aat Asn	aat Asn	tcc Ser 150	cac His	att Ile	ctt Leu	tta Leu	904
agt Ser 155	gct Ala	tta Leu	aaa Lys	tat Tyr	cct Pro 160	gca Ala	aat Asn	gta Val	gca Ala	gtt Val 165	gat Asp	cca Pro	gta Val	gaa Glu	agg Arg 170	952
ttt Phe	ata Ile	ttt Phe	tgg Trp	tct Ser 175	tca Ser	gag Glu	gtg Val	gct Ala	gga Gly 180	agc Ser	ctt Leu	tat Tyr	aga 'Arg	gca Ala 185	gat Asp	1000
ctc Leu	gat Asp	ggt Gly	gtg Val 190	gga Gly	gtg Val	aag Lys	gct Ala	ctg Leu 195	ttg Leu	gag Glu	aca Thr	tca Ser	gag Glu 200	aaa Lys	ata Ile	1048
aca Thr	gct Ala	gtg Val 205	tca Ser	ttg Leu	gat Asp	gtg Val	ctt Leu 210	gat Asp	aag Lys	cgg Arg	ctg Leu	ttt Phe 215	tgg Trp	att Ile	cag Gln	1096
tac Tyr	aac Asn 220	aga Arg	gaa Glu	gga Gly	agc Ser	aat Asn 225	tct Ser	ctt Leu	att Ile	tgc Cys	tcc Ser 230	tgt Cys	gat Asp	tat Tyr	gat Asp	1144
gga Gly 235	ggt Gly	tct Ser	gtc Val	cac His	att Ile 240	agt Ser	aaa Lys	cat His	cca Pro	aca Thr 245	cag Gln	cat His	aat Asn	ttg Leu	ttt Phe 250	1192
gca Ala	atg Met	tcc Ser	ctt Leu	ttt Phe 255	ggt Gly	gac Asp	cgt Arg	atc Ile	ttc Phe 260	tat Tyr	tca Ser	aca Thr	tgg Trp	aaa Lys 265	atg Met	1240
aag Lys	aca Thr	att Ile	tgg Trp 270	ata Ile	gcc Ala	aac Asn	aaa Lys	cac His 275	act Thr	gga Gly	aag Lys	gac Asp	atg Met 280	gtt Val <sup>.</sup>	aga Arg	1288
att	aac	ctc	cat	tca	tca	ttt	gta	cca	ctt	ggt	gaa	ctg	aaa	gta	gtg	1336

Ile	Asn	Leu 285	His	Ser	Ser	Phe	Val 290	Pro	Leu	Gly	Glu	Leu 295	Lys	Val	Val	
cat His	cca Pro 300	ctt Leu	gca Ala	caa Gln	ccc Pro	aag Lys 305	gca Ala	gaa Glu	gat Asp	gac Asp	act Thr 310	tgg Trp	gag Glu	cct Pro	gag Glu	1384
	aaa Lys															1432
gjà aaa	caa Gln	gac Asp	ctc Leu	cag Gln 335	tca Ser	cac His	ttg Leu	tgc Cys	atg Met 340	tgt Cys	gca Ala	gag Glu	gga Gly	tac Tyr 345	gcc Ala	1480
cta Leu	agt Ser	cga Arg	gac Asp 350	cgg Arg	aag Lys	tac Tyr	tgt Cys	gaa Glu 355	gat Asp	gtt Val	aat Asn	gaa Glu	tgt Cys 360	gct Ala	ttt' Phe	1528
tgg Trp	aat Asn	cat His 365	ggc Gly	tgt Cys	act Thr	ctt Leu	999 Gly 370	tgt Cys	aaa Lys	aac Asn	acc Thr	cct Pro 375	gga Gly	tcc Ser	tat Tyr	1576
tac Tyr	tgc Cys 380	acg Thr	tgc Cys	cct Pro	gta Val	gga Gly 385	ttt Phe	gtt Val	ctg Leu	ctt Leu	cct Pro 390	gat Asp	Gly aaa	aaa Lys	cga Arg	1624
	cat His															1672
	tgt Cys															1720
	gtg Val															1768
	aat Asn															1816
	gaa Glu 460															1864
	tgt Cys															1912
caa Gln	gat Asp	att Ile	cga Arg	cac His 495	atg Met	cat His	ttt Phe	gat Asp	gga Gly 500	aca Thr	gac Asp	tat Tyr	gga Gly	act Thr 505	ctg Leu	1960
ctc Leu	agc Ser	cag Gln	cag Gln 510	atg Met	gga Gly	atg Met	gtt Val	tat Tyr 515	gcc Ala	cta Leu	gat Asp	cat His	gac Asp 520	cct Pro	gtg Val	2008
gaa Glu	aat Asn	aag Lys	ata Ile	tac Tyr	ttt Phe	gcc Ala	cat His	aca Thr	gcc Ala	ctg Leu	aag Lys	tgg Trp	ata Ile	gag Glu	aga Arg	2056

525 530 535

gct Ala	aat Asr 540	r Met	g gat : Asj	gg Gl	t tc y Se:	c cag r Gl: 54!	n Arg	a gaa g Gli	a ag u Ar	g ct g Le	t at u Il 55	e Glu	g gaa 1 Gl	a gg u Gl	a gta y Val	2104
555	val	. Pro	) GI	1 GI	56) Б	ı Ala	a Val	L Ası	o Tr	p Il 56	e Gl: 5	y Arg	y Arg	g Phe	tat Tyr 570	2152
-+1	****	. Asp	ALC	575	2 V TIŽE	s sei	г цес	r 116	580 580	y Arq O	g Sei	r Asp	Let	1 Ası 589	_	2200
aaa Lys	cgt Arg	tcc Ser	aaa Lys 590	, 116	a ato e Ile	act Thi	aag Lys	gag Glu 595	ı Ası	c ato	c tct e Sei	c caa c Gln	cca Pro 600	Arg	a gga g Gly	2248
att Ile	gct Ala	gtt Val 605	HIS	cca Pro	ato Met	gco Ala	aag Lys 610	Arc	tta Lei	a tto 1 Phe	tgg Trp	act Thr 615	Asp	aca Thr	Gly ggg	2296
att Ile	aat Asn 620	cca Pro	cga Arg	att	gaa Glu	agt Ser 625	Ser	tcc Ser	ter	caa Glr	Gly 630	r Leu	ggc	cgt Arg	ctg Leu	2344
gtt Val 635	ata Ile	gcc Ala	agc Ser	tct Ser	gat Asp 640	Leu	atc Ile	tgg Trp	Pro	agt Ser 645	Gly	ata Ile	acg Thr	att Ile	gac Asp 650	2392
ttc Phe	tta Leu	act Thr	gac Asp	aag Lys 655	ttg Leu	tac Tyr	tgg Trp	tgc Cys	gat Asp 660	Ala	aag Lys	cag Gln	tct Ser	gtg Val 665	att Ile	2440
gaa Glu	atg Met	gcc Ala	aat Asn 670	ctg Leu	gat Asp	ggt Gly	tca Ser	aaa Lys 675	cgc Arg	cga Arg	aga Arg	ctt Leu	acc Thr 680	cag Gln	aat Asn	2488
gat Asp	gta Val	ggt Gly 685	cac His	cca Pro	ttt Phe	gct Ala	gta Val 690	gca Ala	gtg Val	ttt Phe	gag Glu	gat Asp 695	tat Tyr	gtg Val	tgg Trp	2536
	tca Ser 700	gat Asp	tgg Trp	gct Ala	atg Met	cca Pro 705	tca Ser	gta Val	ata Ile	aga Arg	gta Val 710	aac Asn	aag Lys	agg Arg	act Thr	2584
ggc Gly 715	aaa Lys	gat Asp	aga Arg	gta Val	cgt Arg 720	ctc Leu	caa Gln	ggc Gly	agc Ser	atg Met 725	ctg Leu	aag Lys	ccc Pro	tca Ser	tca Ser 730	2632
ctg ( Leu '	gtt Val	gtg Val	gtt Val	cat His 735	cca Pro	ttg Leu	gca Ala	aaa Lys	cca Pro 740	gga Gly	gca Ala	gat Asp	ccc Pro	tgc Cys 745	tta Leu	2680
tat (	caa Gln .	ASI	gga Gly 750	ggc	tgt Cys	gaa Glu	His	att Ile 755	tgc Cys	aaa Lys	aag Lys	Arg	ctt Leu 760	gga Gly	act Thr	2728
gct t Ala 1	rīb (	tgt Cys : 765	tcg Ser	tgt Cys	cgt Arg	Glu	ggt Gly 770	ttt Phe	atg Met	aaa Lys	Ala	tca Ser 775	gat Asp	gly aaa	aaa Lys	2776

acg Thr	tgt Cys 780	Let	g gct L Ala	ctg Leu	gat Asp	ggt Gly 785	His	cag Gln	ctg Leu	ttg Lev	g gca 1 Ala 790	Gly	ggt Gly	gaa Glu	gtt Val	2824
gat Asp 795	Leu	aag Lys	aac Asn	caa Gln	gta Val 800	Thr	cca Pro	ttg Leu	gac Asp	ato Ile 805	Leu	tcc Ser	aag	act Thr	aga Arg 810	2872
gtg Val	tca Ser	gaa Glu	gat Asp	aac Asn 815	Ile	aca Thr	gaa Glu	tct Ser	caa Gln 820	His	atg Met	cta Leu	gtg Val	gct Ala 825	gaa Glu	2920
atc Ile	atg Met	gtg Val	ser 830	Asp	caa Gln	gat Asp	gac Asp	tgt Cys 835	gct Ala	cct	gtg Val	gga Gly	tgc Cys 840	Ser	atg Met	2968
tat Tyr	gct Ala	cgg Arg 845	Cys	att Ile	tca Ser	gag Glu	gga Gly 850	gag Glu	gat Asp	gcc	aca Thr	tgt Cys 855	Gln	tgt Cys	ttg Leu	3016
aaa Lys	gga Gly 860	Phe	gct Ala	Gly aaa	gat Asp	gga Gly 865	aaa Lys	cta Leu	tgt Cys	tct Ser	gat Asp 870	ata Ile	gat Asp	gaa Glu	tgt Cys	3064
gag Glu 875	Met	ggt Gly	gtc Val	cca Pro	gtg Val 880	tgc Cys	ccc Pro	cct Pro	gcc Ala	tcc Ser 885	tcc Ser	aag Lys	tgc Cys	atc Ile	aac Asn 890	3112
acc Thr	gaa Glu	ggt Gly	ggt Gly	tat Tyr 895	gtc Val	tgc Cys	cgg Arg	tgc Cys	tca Ser 900	gaa Glu	ggc	tac Tyr	caa Gln	gga Gly 905	gat Asp	3160
ej aaa	att Ile	cac His	tgt Cys 910	ctt Leu	gat Asp	att Ile	gat Asp	gag Glu 915	tgc Cys	caa Gln	ctg Leu	Gly	gtg Val 920	cac His	agc Ser	3208
tgt Cys	gga Gly	gag Glu 925	aat Asn	gcc Ala	agc Ser	tgc Cys	aca Thr 930	aat Asn	aca Thr	gag Glu	gga Gly	ggc Gly 935	tat Tyr	acc Thr	tgc Cys	3256
atg Met	tgt Cys 940	gct Ala	gga Gly	cgc Arg	ctg Leu	tct Ser 945	gaa Glu	cca Pro	gga Gly	ctg Leu	att Ile 950	tgc Cys	cct Pro	gac Asp	tct Ser	3304
act Thr 955	cca Pro	ccc Pro	cct Pro	cac His	ctc Leu 960	agg Arg	gaa Glu	gat Asp	gac Asp	cac His 965	cac His	tat Tyr	tcc Ser	gta Val	aga Arg 970	3352
aat Asn	agt Ser	gac Asp	tct Ser	gaa Glu 975	tgt Cys	ccc Pro	ctg Leu	tcc Ser	cac His 980	gat Asp	gly aaa	tac Tyr	tgc Cys	ctc Leu 985	cat His	3400
Asp	Gly	Val	Cys 990	Met	Tyr	Ile	Glu	Ala 995	Leu	Asp	aag Lys	Tyr 1	Ala 000	Cys	Asn	3448
tgt Cys	Val	gtt Val 005	ggc	tac Tyr	atc Ile	Gly	gag Glu 010	cga Arg	tgt Cys	cag Gln	tac Tyr 1	cga Arg 015	gac Asp	ctg. Leu	aag Lys	3496

tgg tgg gaa ctg cgc cac gct ggc cac ggg cag cag cag aag gtc atc Trp Trp Glu Leu Arg His Ala Gly His Gly Gln Gln Gln Lys Val Ile 1020 1025 1030	3544
gtg gtg gct gtc tgc gtg gtg gtg ctt gtc atg ctg ctc ctc ctg agc Val Val Ala Val Cys Val Val Val Leu Val Met Leu Leu Leu Leu Ser 1035 1040 1045 1050	3592
ctg tgg ggg gcc cac tac tac agg act cag aag ctg cta tcg aaa aac Leu Trp Gly Ala His Tyr Tyr Arg Thr Gln Lys Leu Leu Ser Lys Asn 1055 1060 1065	3640
cca aag aat cct tat gag gag tcg agc aga gat gtg agg agt cgc agg Pro Lys Asn Pro Tyr Glu Glu Ser Ser Arg Asp Val Arg Ser Arg Arg 1070 1075 1080	3688
cct gct gac act gag gat ggg atg tcc tct tgc cct caa cct tgg ttt Pro Ala Asp Thr Glu Asp Gly Met Ser Ser Cys Pro Gln Pro Trp Phe 1085 1090 1095	3736
gtg gtt ata aaa gaa cac caa gac ctc aag aat ggg ggt caa cca gtg Val Val Ile Lys Glu His Gln Asp Leu Lys Asn Gly Gly Gln Pro Val 1100 1105 1110	3784
gct ggt gag gat ggc cag gca gca gat ggg tca atg caa cca act tca Ala Gly Glu Asp Gly Gln Ala Ala Asp Gly Ser Met Gln Pro Thr Ser 1115 1120 1125 1130	3832
tgg agg cag gag ccc cag tta tgt gga atg ggc aca gag caa ggc tgc Trp Arg Gln Glu Pro Gln Leu Cys Gly Met Gly Thr Glu Gln Gly Cys 1135 1140 1145	3880
tgg att cca gta tcc agt gat aag ggc tcc tgt ccc cag gta atg gag Trp Ile Pro Val Ser Ser Asp Lys Gly Ser Cys Pro Gln Val Met Glu 1150 1155 1160	3928
cga agc ttt cat atg ccc tcc tat ggg aca cag acc ctt gaa ggg ggt Arg Ser Phe His Met Pro Ser Tyr Gly Thr Gln Thr Leu Glu Gly Gly 1165 1170 1175	3976
gtc gag aag ccc cat tct ctc cta tca gct aac cca tta tgg caa caa Val Glu Lys Pro His Ser Leu Leu Ser Ala Asn Pro Leu Trp Gln Gln 1180 1185 1190	4024
agg gcc ctg gac cca cca cac caa atg gag ctg act cag tga Arg Ala Leu Asp Pro Pro His Gln Met Glu Leu Thr Gln 1195 1200 1205	4066
aaactggaat taaaaggaaa gtcaagaaga atgaactatg tcgatgcaca gtatcttttc	4126
tttcaaaagt agagcaaaac tataggtttt ggttccacaa tctctacgac taatcaccta	4186
ctcaatgcct ggagacagat acgtagttgt gcttttgttt gctcttttaa gcagtctcac	4246
tgcagtctta tttccaagta agagtactgg gagaatcact aggtaactta ttagaaaccc	4306
aaattgggac aacagtgctt tgtaaattgt gttgtcttca gcagtcaata caaatagatt	4366
tttgtttttg ttgttcctgc agccccagaa gaaattaggg gttaaagcag acagtcacac	4426

tggtttggtc agttacaaag taatttettt gatetggaca gaacatttat atcagtttca 4486 tgaaatgatt ggaatattac aataccgtta agatacagtg taggcattta actcctcatt 4546 ggcgtggtcc atgctgatga ttttgccaaa atgagttgtg atgaatcaat gaaaaatgta 4606 atttagaaac tgatttcttc agaattagat ggccttattt tttaaaaatat ttgaatgaaa 4666 acattttatt tttaaaatat tacacaggag geetteggag tttettagte attactgtee 4726 ttttccccta cagaattttc cctcttggtg tgattgcaca gaatttgtat gtattttcag 4786 ttacaagatt gtaagtaaat tgcctgattt gttttcatta tagacaacga tgaatttctt 4846 ctaattattt aaataaaatc accaaaaaca t 4877

<210> 26

<211> 1207

<212> PRT

<213> Homo sapiens

<223> Human EGF

<400> 26

Met Leu Leu Thr Leu Ile Ile Leu Leu Pro Val Val Ser Lys Phe Ser 10 Phe Val Ser Leu Ser Ala Pro Gln His Trp Ser Cys Pro Glu Gly Thr 20 25 Leu Ala Gly Asn Gly Asn Ser Thr Cys Val Gly Pro Ala Pro Phe Leu 40 Ile Phe Ser His Gly Asn Ser Ile Phe Arg Ile Asp Thr Glu Gly Thr 60 Asn Tyr Glu Gln Leu Val Val Asp Ala Gly Val Ser Val Ile Met Asp 70 75 Phe His Tyr Asn Glu Lys Arg Ile Tyr Trp Val Asp Leu Glu Arg Gln 90 Leu Leu Gln Arg Val Phe Leu Asn Gly Ser Arg Gln Glu Arg Val Cys 105 110 Asn Ile Glu Lys Asn Val Ser Gly Met Ala Ile Asn Trp Ile Asn Glu 120 Glu Val Ile Trp Ser Asn Gln Glu Gly Ile Ile Thr Val Thr Asp 135 Met Lys Gly Asn Asn Ser His Ile Leu Leu Ser Ala Leu Lys Tyr Pro 150 155 Ala Asn Val Ala Val Asp Pro Val Glu Arg Phe Ile Phe Trp Ser Ser 170 Glu Val Ala Gly Ser Leu Tyr Arg Ala Asp Leu Asp Gly Val Gly Val Lys Ala Leu Leu Glu Thr Ser Glu Lys Ile Thr Ala Val Ser Leu Asp Val Leu Asp Lys Arg Leu Phe Trp Ile Gln Tyr Asn Arg Glu Gly Ser 215 220 Asn Ser Leu Ile Cys Ser Cys Asp Tyr Asp Gly Gly Ser Val His Ile 235 Ser Lys His Pro Thr Gln His Asn Leu Phe Ala Met Ser Leu Phe Gly 245 250 Asp Arg Ile Phe Tyr Ser Thr Trp Lys Met Lys Thr Ile Trp Ile Ala 265 Asn Lys His Thr Gly Lys Asp Met Val Arg Ile Asn Leu His Ser Ser 280

```
Phe Val Pro Leu Gly Glu Leu Lys Val Val His Pro Leu Ala Gln Pro
                         295
 Lys Ala Glu Asp Asp Thr Trp Glu Pro Glu Gln Lys Leu Cys Lys Leu
                     310
                                         315
 Arg Lys Gly Asn Cys Ser Ser Thr Val Cys Gly Gln Asp Leu Gln Ser
                 325
                                     330
 His Leu Cys Met Cys Ala Glu Gly Tyr Ala Leu Ser Arg Asp Arg Lys
                                 345
 Tyr Cys Glu Asp Val Asn Glu Cys Ala Phe Trp Asn His Gly Cys Thr
                             360
 Leu Gly Cys Lys Asn Thr Pro Gly Ser Tyr Tyr Cys Thr Cys Pro Val
                        375
 Gly Phe Val Leu Leu Pro Asp Gly Lys Arg Cys His Gln Leu Val Ser
                    390
                                         395
 Cys Pro Arg Asn Val Ser Glu Cys Ser His Asp Cys Val Leu Thr Ser
                 405
                                     410
 Glu Gly Pro Leu Cys Phe Cys Pro Glu Gly Ser Val Leu Glu Arg Asp
             420
                                 425
                                       .
 Gly Lys Thr Cys Ser Gly Cys Ser Ser Pro Asp Asn Gly Gly Cys Ser
         435
                             440
 Gln Leu Cys Val Pro Leu Ser Pro Val Ser Trp Glu Cys Asp Cys Phe
                         455
                                             460
 Pro Gly Tyr Asp Leu Gln Leu Asp Glu Lys Ser Cys Ala Ala Ser Gly
                     470
                                         475
 Pro Gln Pro Phe Leu Leu Phe Ala Asn Ser Gln Asp Ile Arg His Met
                 485
                                     490
His Phe Asp Gly Thr Asp Tyr Gly Thr Leu Leu Ser Gln Gln Met Gly
                                 505
Met Val Tyr Ala Leu Asp His Asp Pro Val Glu Asn Lys Ile Tyr Phe
                             520
                                                 525
Ala His Thr Ala Leu Lys Trp Ile Glu Arg Ala Asn Met Asp Gly Ser
                        535
Gln Arg Glu Arg Leu Ile Glu Glu Gly Val Asp Val Pro Glu Gly Leu
                                         555
Ala Val Asp Trp Ile Gly Arg Arg Phe Tyr Trp Thr Asp Arg Gly Lys
                                    570
Ser Leu Ile Gly Arg Ser Asp Leu Asn Gly Lys Arg Ser Lys Ile Ile
                                585
Thr Lys Glu Asn Ile Ser Gln Pro Arg Gly Ile Ala Val His Pro Met
                            600
Ala Lys Arg Leu Phe Trp Thr Asp Thr Gly Ile Asn Pro Arg Ile Glu
                        615
Ser Ser Ser Leu Gln Gly Leu Gly Arg Leu Val Ile Ala Ser Ser Asp
                    630
                                        635
Leu Ile Trp Pro Ser Gly Ile Thr Ile Asp Phe Leu Thr Asp Lys Leu
                                    650
Tyr Trp Cys Asp Ala Lys Gln Ser Val Ile Glu Met Ala Asn Leu Asp
                                665
Gly Ser Lys Arg Arg Leu Thr Gln Asn Asp Val Gly His Pro Phe
        675
                            680
Ala Val Ala Val Phe Glu Asp Tyr Val Trp Phe Ser Asp Trp Ala Met
                        695
                                            700
Pro Ser Val Ile Arg Val Asn Lys Arg Thr Gly Lys Asp Arg Val Arg
                    710
                                        715
Leu Gln Gly Ser Met Leu Lys Pro Ser Ser Leu Val Val His Pro
                                    730
Leu Ala Lys Pro Gly Ala Asp Pro Cys Leu Tyr Gln Asn Gly Gly Cys
                                745
                                                    750
Glu His Ile Cys Lys Lys Arg Leu Gly Thr Ala Trp Cys Ser Cys Arg
                            760
Glu Gly Phe Met Lys Ala Ser Asp Gly Lys Thr Cys Leu Ala Leu Asp
```

775 780 Gly His Gln Leu Leu Ala Gly Gly Glu Val Asp Leu Lys Asn Gln Val 790 795 Thr Pro Leu Asp Ile Leu Ser Lys Thr Arg Val Ser Glu Asp Asn Ile 810 Thr Glu Ser Gln His Met Leu Val Ala Glu Ile Met Val Ser Asp Gln 820 825 Asp Asp Cys Ala Pro Val Gly Cys Ser Met Tyr Ala Arg Cys Ile Ser 835 840 Glu Gly Glu Asp Ala Thr Cys Gln Cys Leu Lys Gly Phe Ala Gly Asp 855 Gly Lys Leu Cys Ser Asp Ile Asp Glu Cys Glu Met Gly Val Pro Val 870 875 Cys Pro Pro Ala Ser Ser Lys Cys Ile Asn Thr Glu Gly Gly Tyr Val 890 Cys Arg Cys Ser Glu Gly Tyr Gln Gly Asp Gly Ile His Cys Leu Asp 905 Ile Asp Glu Cys Gln Leu Gly Val His Ser Cys Gly Glu Asn Ala Ser 920 Cys Thr Asn Thr Glu Gly Gly Tyr Thr Cys Met Cys Ala Gly Arg Leu 940 935 Ser Glu Pro Gly Leu Ile Cys Pro Asp Ser Thr Pro Pro Pro His Leu 950 · 955 Arg Glu Asp Asp His His Tyr Ser Val Arg Asn Ser Asp Ser Glu Cys 965 970 Pro Leu Ser His Asp Gly Tyr Cys Leu His Asp Gly Val Cys Met Tyr 980 985 Ile Glu Ala Leu Asp Lys Tyr Ala Cys Asn Cys Val Val Gly Tyr Ile 995 1000 1005 Gly Glu Arg Cys Gln Tyr Arg Asp Leu Lys Trp Trp Glu Leu Arg His 1010 1015 1020 Ala Gly His Gly Gln Gln Lys Val Ile Val Val Ala Val Cys Val 1030 1035 Val Val Leu Val Met Leu Leu Leu Ser Leu Trp Gly Ala His Tyr 1045 1050 1055 Tyr Arg Thr Gln Lys Leu Leu Ser Lys Asn Pro Lys Asn Pro Tyr Glu 1060 1065 1070 Glu Ser Ser Arg Asp Val Arg Ser Arg Arg Pro Ala Asp Thr Glu Asp 1075 1080 1085 Gly Met Ser Ser Cys Pro Gln Pro Trp Phe Val Val Ile Lys Glu His 1095 1100 Gln Asp Leu Lys Asn Gly Gly Gln Pro Val Ala Gly Glu Asp Gly Gln 1105 1110 1115 Ala Ala Asp Gly Ser Met Gln Pro Thr Ser Trp Arg Gln Glu Pro Gln 1125 1130 Leu Cys Gly Met Gly Thr Glu Gln Gly Cys Trp Ile Pro Val Ser Ser 1140 1145 1150 Asp Lys Gly Ser Cys Pro Gln Val Met Glu Arg Ser Phe His Met Pro 1155 1160 Ser Tyr Gly Thr Gln Thr Leu Glu Gly Gly Val Glu Lys Pro His Ser 1175 1180 Leu Leu Ser Ala Asn Pro Leu Trp Gln Gln Arg Ala Leu Asp Pro Pro 1190 1195 His Gln Met Glu Leu Thr Gln 1205

<210> 27 <211> 4749

<212> DNA

<213> Mus musculus <220> <221> CDS <222> (354)..(4007) <220> <223> Murine EGF <400> 27 aaaaaaggag aagggattcc tatctgtata tagggaagga atcctatctg catatttcgt 60 tgttagcacc atccctcatc ccggtgggct tggaactttc catcaattct ttcctgtctc 120 gtttctcttt catcctttgc ctggttgtgc ctgtctcagg gagaaatcag tcacctgcag 180 gccttgcagg gctcttaggc tctgggaaat ttgtcatacg ggtgtcaggt acttcttatt 240 gctgtccaaa gggaaaaaaa aagtgagaca aagaactctc ccggagcctt tccggctgca 300 ctcagaggct ctcgagaggt gcagaaggac ctggaaaggc agctaaataa aag atg ecc tgg gge ega agg cea acc tgg etg ttg etc gec tte etg etg gtg 404 Pro Trp Gly Arg Arg Pro Thr Trp Leu Leu Leu Ala Phe Leu Leu Val ttt tta aag att agc ata ctc agc gtc aca gca tgg cag acc ggg aac 452 Phe Leu Lys Ile Ser Ile Leu Ser Val Thr Ala Trp Gln Thr Gly Asn 20 tgt cag cca ggt cct ctc gag aga agc gag aga agc ggg act tgt gcc 500 Cys Gln Pro Gly Pro Leu Glu Arg Ser Glu Arg Ser Gly Thr Cys Ala ggt cct gcc ccc ttc cta gtt ttc tca caa gga aag agc atc tct cgg Gly Pro Ala Pro Phe Leu Val Phe Ser Gln Gly Lys Ser Ile Ser Arg 50 55 att gac cca gat gga aca aat cac cag caa ttg gtg gtg gat gct ggc 596 Ile Asp Pro Asp Gly Thr Asn His Gln Gln Leu Val Val Asp Ala Gly atc tca gca gac atg gat att cat tat aaa aaa gag aga ctc tat tgg 644 Ile Ser Ala Asp Met Asp Ile His Tyr Lys Lys Glu Arg Leu Tyr Trp 85 gtg gat gta gaa aga caa gtt ttg cta aga gtt ttc ctt aac ggg aca 692 Val Asp Val Glu Arg Gln Val Leu Leu Arg Val Phe Leu Asn Gly Thr gga cta gag aaa gtg tgc aat gta gag agg aag gtg tct ggg ctg gcc Gly Leu Glu Lys Val Cys Asn Val Glu Arg Lys Val Ser Gly Leu Ala ata gac tgg ata gat gat gaa gtt ctc tgg gta gac caa cag aac gga Ile Asp Trp Ile Asp Asp Glu Val Leu Trp Val Asp Gln Gln Asn Gly 140

val	L 11	e Th	r va.	1 Th	r As <sub>l</sub> O	p Met	: Thi	r Gly	7 Ly:	s As: 5	n Se	r Aro	y Va:	l Le 16		836
agt Ser	t too	c tta	a aaa u Ly: 16!	s Hi	t ccg	g tca Ser	a aat : Asr	ata 1 Ile 170	e Ala	a gtg a Val	g ga l As	t cca p Pro	ata Ile 175	e Gl	g agg u Arg	884
ttg Leu	g ato Met	tti Phe 180	s III	g tot p Sei	tca Sex	a gag Glu	gtg Val 185	. Thr	. Gl <sup>2</sup> s aad	z ago 7 Sei	c Lei	t cac u His 190	Arc	gca J Ala	a cac a His	932
ctc Leu	aaa Lys 195	s GT	gtt Val	gat L Asp	gta Val	a aaa Lys 200	Thr	ctg Leu	cto Lei	g gag ı Glu	g aca 1 Thi 205	c Gly	gga Gly	ata Ile	tcg Ser	980
gtg Val 210	Let	act Thr	cto Lev	g gat 1 Asp	gto Val 215	. Leu	gac Asp	aaa Lys	. cgg	g cto Lev 220	Phe	tgg Trp	gtt Val	Glr	gac Asp 225	1028
ber	GIY	GIU	. сту	230	Hls		Tyr	Ile	His 235	Ser	Cys	gat Asp	Tyr	Glu 240	Gly	1076
CLY	DCI	Val	245	neu	тте	Arg	HIS	GIn 250	Ala	Arg	His	agt Ser	Leu 255	Ser	Ser	1124
atg Met	gcc Ala	ttt Phe 260	ttt Phe	ggt	gat Asp	cgg Arg	atc Ile 265	ttc Phe	tac Tyr	tca Ser	gtg Val	ttg Leu 270	aaa Lys	agc Ser	aag Lys	1172
nia	275	TTP	116	ALA	ASI	ьуs 280	HIS	Thr	Gly	Lys	Asp 285	acg Thr	Val	Arg	Ile	1220
aac Asn 290	ctc Leu	cat His	cca Pro	tcc Ser	ttt Phe 295	gtg Val	aca Thr	cct Pro	gga Gly	aaa Lys 300	ctg Leu	atg Met	gta Val	gta Val	cac His 305	1268
cct Pro	cgt Arg	gca Ala	cag Gln	ccc Pro 310	agg Arg	aca Thr	gag Glu	gac Asp	gct Ala 315	gct Ala	aag Lys	gat Asp	cct Pro	gac Asp 320	ccc Pro	1316
GIU	ren	ьeu	цуs 325	GIN	Arg	GIÀ	Arg	Pro 330	Cys	Arg	Phe	ggt Gly	Leu 335	Cys	Glu	1364
cga Arg	gac Asp	ccc Pro 340	aag Lys	tcc Ser	cac His	ser	agc Ser 345	gca Ala	tgc Cys	gct Ala	gag Glu	ggc Gly 350	tac Tyr	acg Thr	tta Leu	1412
ser 1	cga Arg 355	gac Asp	cgg Arg	aag Lys	Tyr	tgc Cys 360	gaa Glu	gat Asp	gtc Val	Asn	gaa Glu 365	tgt Cys	gcc Ala	act Thr	cag Gln	1460
aat o Asn I 370	cac His	ggc	tgt Cys	TIIL	ctt Leu 375	Gly (	tgt ( Cys (	gaa : Glu :	Asn	acc Thr 380	cct Pro	gga ( Gly (	tcc (	Iyr	cac His 385	1508

				aca Thr 390												1556
				tcc Ser												1604
_	_	Leu		tca Ser	-				_				_			1652
			_	gat Asp		_		_			_		_		_	1700
				agc Ser												1748
				ttt Phe 470												1796
				gga Gly												1844
				atg Met												1892
				gga Gly												1940
				ttt Phe												1988
			_	tcc Ser 550	-					_			_ =		_	2036
				ctt Leu												2084
				aag Lys												2132
				atc Ile												2180
				agg Arg												2228
tct	cca	cgg	att	gaa	agc	gct	tcc	ctt	caa	ggt	tcc	gac	cgg	gtg	ctg	2276

Ser	Pro	Arg	Ile	Glu 630		Ala	Ser	Leu	Gln 635		Ser	Asp	Arg	Val 640	Leu	
ata Ile	gcc Ala	agc Ser	tcc Ser 645	Asn	cta Leu	ctg Leu	gaa Glu	Pro 650	agt Ser	gga Gly	atc Ile	acg Thr	att Ile 655	gac Asp	tac Tyr	2324
tta Leu	aca Thr	gac Asp 660	act Thr	ttg Leu	tac Tyr	tgg Trp	tgt Cys 665	gac Asp	acc Thr	aag Lys	agg Arg	tct Ser 670	Val	att Ile	gaa Glu	2372
atg Met	gcc Ala 675	aat Asn	ctg Leu	gat Asp	gly	tcc Ser 680	aaa Lys	cgc Ārg	cga Arg	aga Arg	ctt Leu 685	atc Ile	cag Gln	aac Asn	gac Asp	2420
gta Val 690	ggt Gly	cac His	ccc Pro	ttc Phe	tct Ser 695	cta Leu	gcc Ala	gtg Val	ttt Phe	gag Glu 700	gat Asp	cac His	ctg Leu	tgg Trp	gtc Val 705	2468
tcg Ser	gat Asp	tgg Trp	gct Ala	atc Ile 710	cca Pro	tcg Ser	gta Val	ata Ile	agg Arg 715	gtg Val	aac Asn	aag Lys	agg Arg	act Thr 720	ggc	2516
caa Gln	aac Asn	agg Arg	gta Val 725	Arg	ctt Leu	caa Gln	ggc	agc Ser 730	atg Met	ctg Leu	aag Lys	ccc Pro	tcg Ser 735	tca Ser	ctg Leu	2564
gtt Val	gtg Val	gtc Val 740	cat His	cca Pro	ttg Leu	gca Ala	aaa Lys 745	cca Pro	ggt Gly	gca Ala	gat Asp	ccc Pro 750	tgc Cys	tta Leu	tac Tyr	2612
agg Arg	aat Asn 755	gga Gly	ggc	tgt Cys	gaa Glu	cac His 760	atc Ile	tgc Cys	caa Gln	gag Glu	agc Ser 765	ctg Leu	Gly	aca Thr	gct Ala	2660
cgg Arg 770	tgt Cys	ttg Leu	tgt Cys	cgt Arg	gaa Glu 775	ggt Gly	ttt Phe	gtg Val	aag Lys	gcc Ala 780	tgg Trp	gat Asp	Gly 999	aaa Lys	atg Met 785	2708
tgt Cys	ctc Leu	cct Pro	cag Gln	gat Asp 790	tat Tyr	cca Pro	atc Ile	ctg Leu	tca Ser 795	ggt Gly	gaa Glu	aat Asn	gct Ala	gat Asp 800	ctt Leu	2756
agt Ser	aaa Lys	gag Glu	gtg Val 805	aca Thr	tca Ser	ctg Leu	agc Ser	aac Asn 810	tcc Ser	act Thr	cag Gln	gct Ala	gaa Glu 815	gta Val	cca Pro	2804
gac Asp	gat Asp	gat Asp 820	Gly aaa	aca Thr	gaa Glu	tct Ser	tcc Ser 825	aca Thr	cta Leu	gtg Val	gct Ala	gaa Glu 830	atc Ile	atg Met	gtg Val	2852
tca Ser	ggc Gly 835	atg Met	aac Asn	tat Tyr	gaa Glu	gat Asp 840	gac Asp	tgt Cys	ggt Gly	ccc Pro	999 Gly 845	gly ggg	tgt Cys	gga Gly	agc Ser	2900
cat His 850	gct Ala	cga Arg	tgc Cys	gtt Val	tca Ser 855	gac Asp	gga Gly	gag Glu	act Thr	gct Ala 860	gag Glu	tgt Cys	cag Gln	Cys	ctg Leu 865	2948
aaa Lys	Gly 999	ttt Phe	gcc Ala	agg Arg	gat Asp	gga Gly	aac Asn	ctg Leu	tgt Cys	tct Ser	gat Asp	ata Ile	gat Asp	gag Glu	tgt Cys	2996

	870		•	
		875	880	
var neu mra r	ga tcg gac tgc rg Ser Asp Cys 85	CCC agc acc tcg Pro Ser Thr Ser 890	tcc agg tgc atc aac Ser Arg Cys Ile Asn 895	3044
act gaa ggt g Thr Glu Gly G 900	TY TYL VAL CYS	aga tgc tca gaa Arg Cys Ser Glu 905	ggc tac gaa gga gac Gly Tyr Glu Gly Asp 910	3092
ggg atc tcc t Gly Ile Ser C 915	gt ttc gat att ys Phe Asp Ile . 920	Asp Gin Cys Gin	cgg ggg gcg cac aac Arg Gly Ala His Asn 925	3140
tgc gct gag a Cys Ala Glu A 930	at gcc gcc tgc a sn Ala Ala Cys 1 935	acc aac acg gag Thr Asn Thr Glu 940	gga ggc tac aac tgc Gly Gly Tyr Asn Cys 945	3188
acc tgc gca g Thr Cys Ala G	gc cgc cca tcc ( ly Arg Pro Ser ( 950	tcg ccc gga cgg : Ser Pro Gly Arg : 955	agt tgc cct gac tct Ser Cys Pro Asp Ser 960	3236
acc gca ccc to Thr Ala Pro Se	er neg neg Gil (	gaa gat ggc cac o Glu Asp Gly His I 970	cat ttg gac cga aat His Leu Asp Arg Asn 975	3284
agt tat cca gg Ser Tyr Pro GJ 980	y cys pro ser s	cca tat gat gga t Ser Tyr Asp Gly 1 985	tac tgc ctc aat ggt Tyr Cys Leu Asn Gly 990	3332
ggc gtg tgc at Gly Val Cys Me 995	g cat att gaa t t His Ile Glu S 1000	er Leu Asp Ser T	ac aca tgc aac tgt Tyr Thr Cys Asn Cys 005	3380
gtt att ggc ta Val Ile Gly Ty 1010	t tot ggg gat o r Ser Gly Asp A 1015	ga tgt cag act c rg Cys Gln Thr A 1020	ga gac cta cga tgg arg Asp Leu Arg Trp 1025	3428
tgg gag ctg cg Trp Glu Leu Ar	t cat get gge t g His Ala Gly T 1030	ac ggg cag aag c yr Gly Gln Lys H 1035	at gac atc atg gtg is Asp Ile Met Val 1040	3476
gtg gct gtc tg Val Ala Val Cy 104	s wer var Ara Le	tg gtc ctg ctg c eu Val Leu Leu L 1050	tc ctc ttg ggg atg eu Leu Leu Gly Met 1055	3524
tgg ggg act tac Trp Gly Thr Ty: 1060	c tac tac agg ac r Tyr Tyr Arg Tl 100	nr Arg Lys Gln Le	ta tca aac ccc cca eu Ser Asn Pro Pro 1070	3572
aag aac cct tgt Lys Asn Pro Cys 1075	gat gag cca ag Asp Glu Pro Se 1080	gc gga agt gtg ag er Gly Ser Val Se 108	gc agc agc ggg ccc er Ser Ser Gly Pro 85	3620
gac agc agc agc Asp Ser Ser Ser 1090	ggg gca gct gt Gly Ala Ala Va 1095	g gct tct tgt co al Ala Ser Cys Pr 1100	cc caa cct tgg ttt co Gln Pro Trp Phe 1105	3668
.ar var hea Gra	, aaa cac caa ga Lys His Gln As 1110	ac ccc aag aat gg p Pro Lys Asn Gl 1115	gg agt ctg cct gcg Y Ser Leu Pro Ala 1120	<b>3716</b> .

gat ggt acg aat ggt gca gta gta gat gct ggc ctg tct ccc tcc ctg 3764 Asp Gly Thr Asn Gly Ala Val Val Asp Ala Gly Leu Ser Pro Ser Leu 1125 1130 1135	
cag ctc ggg tca gtg cat ctg act tca tgg aga cag aag ccc cac ata 3812 Gln Leu Gly Ser Val His Leu Thr Ser Trp Arg Gln Lys Pro His Ile 1140 1145 1150	
gat gga atg ggc aca ggg caa agc tgc tgg att cca cca tca agt gac 3860 Asp Gly Met Gly Thr Gly Gln Ser Cys Trp Ile Pro Pro Ser Ser Asp 1155 1160 1165	
aga gga ccc cag gaa ata gag gga aac tcc cac cta ccc tcc tac aga 3908 Arg Gly Pro Gln Glu Ile Glu Gly Asn Ser His Leu Pro Ser Tyr Arg 1170 1185	
cct gtg ggg ccg gag aag ctg cat tct ctc cag tca gct aat gga tcg 3956 Pro Val Gly Pro Glu Lys Leu His Ser Leu Gln Ser Ala Asn Gly Ser 1190 1195 1200	
tgt cac gaa agg gct cca gac ctg cca cgg cag aca gag cca gtt aag 4004 Cys His Glu Arg Ala Pro Asp Leu Pro Arg Gln Thr Glu Pro Val Lys 1205 1210 1215	
tag aaactgggag tagacagaag gtacagaagg gaaaataaca aaccaggctg 4057	
atgatggtag agtgctacag acttggtact ccagtttcca cggctaatca ctgctcgctc 4117	
agggteetga agatagetge acagetgeag agetgeacag egggataget gegaettttg 4177	
cttcttgctt taagcagttc cactgaagat actcaaaaga gaagtggaga aaatcattag 4237	
aaaccaaagt caagacatte atatataage tgtgtettet teactggaeg gtttgeetet 4297	
tttccttttg cctcagaagg agtgggttaa agcaggtgac cccatgctct gtcaacccct 4357	
gaataaatga tgtgatctac atagaagtct tagctcactc tcaggaacgc ttggaacact 4417	
ataacttttg ctatgatata ctgccaagtg tggcccatgc tcataattgt gccttctgaa 4477	
ttgtgataaa ttagtgaaaa aactgtaact tagaatctga tttattcagg attagatcat 4537	
ctttttatac tataaaaatc ttcgaatgaa aatatttaac tttaaaaaca ttaccttaat 4597	
cattgtcttt tcttcttgaa gtctttccca gtgaaaacgc tcaattctgc tgtttccata 4657	
gaatttttaa tttattttaa gacatgagat tgtaaacaaa ttgcttgatt tattttatcc 4717	
taattattta aataaaatca ccctaaagca tc 4749	

<210> 28

<211> 1217

<212> PRT

<213> Mus musculus

<223> Murine EGF

<400> 28

Met Pro Trp Gly Arg Arg Pro Thr Trp Leu Leu Leu Ala Phe Leu Leu

1				5					10					15	
Val	Phe	Leu	Lys 20	Ile	Ser	Ile	Leu	Ser 25	Val	Thr	Ala	Trp	Gln 30	Thr	Gly
	_	35		_		Leu	40					45	_		_
	50					Leu 55					60	_			
Arg 65	Ile	Asp	Pro	Asp	Gly 70	Thr	Asn	His	Gln	Gln 75	Leu	Val	Val	Asp	Ala 80
_				85		Asp			90	_	_		_	95	-
			100			Gln		105					110		_
		115		_		Cys	120			_	_	125		_	
	130					Asp 135				-	140				
145					150	Asp			_	155					160
				165		Pro			170					175	
			180			Ser		185		_			190	_	
		195				Val	200					205	_	-	
	210					Val 215					220		_		
225					230	His				235					240
				245		Ile			250					255	
			260			Asp		265					270		
		275				Asn	280			_	_	285			
	290					Phe 295				_	300				
305					310	Arg				315					320
				325		Arg	_	_	330	_	_		_	335	_
	_	_	340			His		345					350		
		355				Tyr	360					365	_		
	370					375 Gly		_			380				_
385					390	Cys				395		_	_	_	400
				405		Asp			410			_	-	415	
			420		•	Gly		425		_			430		
		435				Gln	440				_	445			
	450					455 Pro					460				
465					470	Pro				475					480
	<b>-</b> 75		734 CA	485	<u>y</u>		-111	-10	490	neu	пеп	FIIE	nta	495	SET

Gln	Asp	Ile	Arg 500	His	Met	His	Phe	Asp 505	Gly	Thr	Asp	Tyr	Lys 510	Val	Leu
Leu	Ser	Arg 515	Gln	Met	Gly	Met	Val 520		Ala	Leu	Asp	Tyr 525	Asp	Pro	Val
Glu	Ser 530		Ile	Tyr	Phe	Ala 535		Thr	Ala	Leu	Lys 540	Trp	Ile	Glu	Arg
Ala 545		Met	Asp	Gly	Ser 550	Gln	Arg	Glu	Arg	Leu 555	Ile	Thr	Glu	Gly	Val 560
	Thr	Leu	Glu	Gly 565	Leu	Ala	Leu	Asp	Trp 570		Gly	Arg	Arg	Ile 575	Tyr
Trp	Thr	Asp	Ser 580	Gly	Lys	Ser	Val	Val 585	Gly	Gly	Ser	Asp	Leu 590	Ser	Gly
ГÀв	His	His 595	Arg	Ile	Ile	Ile	Gln 600	Glu	Arg	Ile	Ser	Arg 605	Pro	Arg	Gly
	610					615					620		Asp		
625					630					635			Asp		640
				645					650				Thr	655	
_			660					665					Ser 670		
		675					680					685	Ile		
-	690					695					700		His		
705		-			710					715			Lys		720
-			_	725	_				730				Pro	735	
			740					745					Pro 750		
-	_	755	_	_			760					765	Leu Asp		
	770	-		_	_	775	_				780		Asn		
785	-				790					795			Ala		800
		-		805					810				Glu	815	
			820	_				825					830		Gly
		835					840					845	Cys		
	850			_		855	_	_			860				Glu
865	-	_			870					875			Arg		880
				885					890					895	Gly
			900					905					910		His
_		915					920					925			Asn
	930					935					940				Asp
945		_			950					955			Leu	٠	960
				965					970					975	Asn

980 985 Gly Gly Val Cys Met His Ile Glu Ser Leu Asp Ser Tyr Thr Cys Asn 995 1000 1005 Cys Val Ile Gly Tyr Ser Gly Asp Arg Cys Gln Thr Arg Asp Leu Arg 1015 1020 Trp Trp Glu Leu Arg His Ala Gly Tyr Gly Gln Lys His Asp Ile Met 1030 1035 Val Val Ala Val Cys Met Val Ala Leu Val Leu Leu Leu Leu Gly 1045 1050 Met Trp Gly Thr Tyr Tyr Arg Thr Arg Lys Gln Leu Ser Asn Pro 1065 Pro Lys Asn Pro Cys Asp Glu Pro Ser Gly Ser Val Ser Ser Ser Gly 1080 Pro Asp Ser Ser Ser Gly Ala Ala Val Ala Ser Cys Pro Gln Pro Trp 1095 1100 Phe Val Val Leu Glu Lys His Gln Asp Pro Lys Asn Gly Ser Leu Pro 1110 1115 Ala Asp Gly Thr Asn Gly Ala Val Val Asp Ala Gly Leu Ser Pro Ser 1125 1130 Leu Gln Leu Gly Ser Val His Leu Thr Ser Trp Arg Gln Lys Pro His 1140 1145 Ile Asp Gly Met Gly Thr Gly Gln Ser Cys Trp Ile Pro Pro Ser Ser 1155 1160 Asp Arg Gly Pro Gln Glu Ile Glu Gly Asn Ser His Leu Pro Ser Tyr 1170 1175 1180 Arg Pro Val Gly Pro Glu Lys Leu His Ser Leu Gln Ser Ala Asn Gly 1190 1195 Ser Cys His Glu Arg Ala Pro Asp Leu Pro Arg Gln Thr Glu Pro Val 1210 Lys <210> 29 <211> 4119 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (32)..(514) <220> <223> Human TGF- alpha <400> 29 ctggagagcc tgctgcccgc ccgcccgtaa a atg gtc ccc tcg gct gga cag Met Val Pro Ser Ala Gly Gln 1 ctc gcc ctg ttc gct ctg ggt att gtg ttg gct gcg tgc cag gcc ttq 100 Leu Ala Leu Phe Ala Leu Gly Ile Val Leu Ala Ala Cys Gln Ala Leu 10 gag aac agc acg tcc ccg ctg agt gca gac ccg ccc gtg gct gca gca Glu Asn Ser Thr Ser Pro Leu Ser Ala Asp Pro Pro Val Ala Ala Ala 25 gtg gtg tcc cat ttt aat gac tgc cca gat tcc cac act cag ttc tgc Val Val Ser His Phe Asn Asp Cys Pro Asp Ser His Thr Gln Phe Cys 40 45

			Thr													
gtc	tgc	cat	tct	<b>9</b> 99	tac	gtt	ggt	gca	cgc	tgt	gag	cat	gcg	gac	ctc	292
Val	Cys	His	Ser 75	Gly	Tyr	Val	Gly	Ala 80	Arg	Сув	Glu	His	Ala 85	Asp	Leu	
			gtg Val													340
gtg Val	gtg Val 105	gtc Val	tcc Ser	atc Ile	gtg Val	gcc Ala 110	ctg Leu	gct Ala	gtc Val	ctt Leu	atc Ile 115	atc Ile	aca Thr	tgt Cys	gtg Val	388
			tgc Cys													436
			cgg													484
			cac His 155						tga	agag	gecea	aga g	ggag	gagti	t.	534
ggcd	aggt	gg (	actgt	ggca	g at	caat	caaaç	g aaa	agget	tct	tcag	ggaca	agc a	actgo	ccagag	594
atgo	ctgg	ggt g	gtgco	acaç	ja co	ette	ctact	tgg	gaato	gtaa	tcac	ectgt	gc a	agcct	tttgt	654
ggg	cetto	caa a	aacto	etgto	a ag	gaact	ccgt	cto	gette	9999	ttat	tcaç	gtg	tgac	tagag	714
aaga	aato	ag (	cggac	cacc	ga tt	tcaa	agact	tgt:	taaa	aaaa	gaad	etgca	aaa q	gagad	ggact	774
ccts	gttca	acc 1	taggt	gagg	jt gt	gtgo	cagca	a gtt	ggtg	gtct	gagt	ccad	cat o	gtgtg	gcagtt	834
gtct	tcts	gcc a	agcca	itgga	it to	cago	gctat	ata	attto	ettt	ttaa	atggg	gee i	accto	cccac	894
aaca	ıgaat	tc 1	tgcco	aaca	ıc aç	gaga	attto	tat	agtt	att	gttt	tctg	gtc a	attt	gcctac	954
tggg	gaag	gaa a	agtga	agga	a ac	ggaaa	actgt	tta	aatat	cac	atga	agad	cac 1	tagct	ttaag	1014
agaa	gcto	gta 1	tcctc	taac	c ac	gaga	actct	caa	accaç	geec	aaca	tctt	cc a	atgga	acacat	1074
gaca	ttga	aag a	accat	ccca	a go	ctato	egcca	a cco	ttgg	gaga	tgat	gtct	ta 1	tttat	tagat	1134
ggat	aato	ggt 1	tttat	tttt	a at	ctct	taag	g tca	aatgt	caaa	aagt	ataa	aaa d	ccct	tcaga	1194
cttc	taca	att a	aatga	itgta	ıt gt	gttg	gctga	cto	gaaaa	agct	atac	tgat	ta d	gaaat	gtctg	1254
															ccaga	
ggct	gggt	ta d	etggt	agga	a ta	aagg	gtago	g ggt	tcag	gaaa	tggt	gcca	tt g	gaago	cacaa	1374
agco	ggta	aa t	tgcct	caat	a cg	jttct	ggga	gaa	aact	tag	caaa	tcca	itc a	agcag	ggatc	1434

tgtcccctct gttggggaga gaggaagagt gtgtgtct acacaggata aacccaatac 1494 atattgtact getcagtgat taaatgggtt cactteeteg tgageeeteg gtaagtatgt 1554 ttagaaatag aacattagcc acgagccata ggcatttcag gccaaatcca tgaaaggggg 1614 accagteatt tatttteeat tttgttgett ggttggtttg ttgetttatt tttaaaagga 1674 gaagtttaac tttgctattt attttcgagc actaggaaaa ctattccagt aattttttt 1734 tecteattte catteaggat geeggettta ttaacaaaaa etetaacaag teaceteeac 1794 tatgtgggtc ttcctttccc ctcaagagaa ggagcaattg ttcccctgac atctgggtcc 1854 atctgaccca tggggcctgc ctgtgagaaa cagtgggtcc cttcaaatac atagtggata 1914 gctcatccct aggaattttc attaaaattt ggaaacagag taatgaagaa ataatatat 1974 aactccttat gtgaggaaat gctactaata tctgaaaagt gaaagatttc tatgtattaa 2034 ctcttaagtg cacctagctt attacatcgt gaaaggtaca tttaaaatat gttaaattgg 2094 cttgaaattt tcagagaatt ttgtcttccc ctaattcttc ttccttggtc tggaagaaca 2154 atttctatga attttctctt tattttttt ttataattca gacaattcta tgacccgtgt 2214 cttcattttt ggcactctta tttaacaatg ccacacctga agcacttgga tctgttcaga 2274 gctgaccccc tagcaacgta gttgacacag ctccaggttt ttaaattact aaaataagtt 2334 caagtttaca teeettggge cagatatgtg ggttgagget tgactgtage atcetgetta 2394 gagaccaatc aatggacact ggtttttaga cctctatcaa tcagtagtta gcatccaaga 2454 gactttgcag aggcgtagga atgaggctgg acagatggcg gaacgagagg ttccctgcga 2514 agacttgaga tttagtgtct gtgaatgttc tagttcctag gtccagcaag tcacacctgc 2574 cagtgccctc atccttatgc ctgtaacaca catgcagtga gaggcctcac atatacgcct 2634 ccctagaagt gccttccaag tcagtccttt ggaaaccagc aggtctgaaa aagaggctgc 2694 atcaatgcaa geetggttgg accattgtee atgeeteagg atagaacage etggettatt 2754 tggggatttt tcttctagaa atcaaatgac tgataagcat tggctccctc tgccatttaa 2814 tggcaatggt agtctttggt tagctgcaaa aatactccat ttcaagttaa aaatgcatct 2874 tetaatecat etetgeaage teeetgtgtt teettgeeet ttagaaaatg aattgtteac 2934 tacaattaga gaatcattta acatcctgac ctggtaagct gccacacac tggcagtggg 2994 gagcatcgct gtttccaatg gctcaggaga caatgaaaag cccccattta aaaaaataac 3054 aaacattttt taaaaggcct ccaatactct tatggagcct ggatttttcc cactgctcta 3114 caggotgtga otttttttaa goatootgao aggaaatgtt ttottotaca tggaaagata 3174 gacagcagcc aaccetgate tggaagacag ggeccegget ggacacacgt ggaaccaage 3234 cagggatggg ctggccattg tgtccccgca ggagagatgg gcagaatggc cctagagttc 3294

```
<210> 30
```

## <400> 30

Met Val Pro Ser Ala Gly Gln Leu Ala Leu Phe Ala Leu Gly Ile Val Leu Ala Ala Cys Gln Ala Leu Glu Asn Ser Thr Ser Pro Leu Ser Ala Asp Pro Pro Val Ala Ala Ala Val Val Ser His Phe Asn Asp Cys Pro 40 Asp Ser His Thr Gln Phe Cys Phe His Gly Thr Cys Arg Phe Leu Val Gln Glu Asp Lys Pro Ala Cys Val Cys His Ser Gly Tyr Val Gly Ala 70 75 Arg Cys Glu His Ala Asp Leu Leu Ala Val Val Ala Ala Ser Gln Lys 90 Lys Gln Ala Ile Thr Ala Leu Val Val Val Ser Ile Val Ala Leu Ala 100 105 Val Leu Ile Ile Thr Cys Val Leu Ile His Cys Cys Gln Val Arg Lys 115 120 His Cys Glu Trp Cys Arg Ala Leu Ile Cys Arg His Glu Lys Pro Ser 135 140 Ala Leu Leu Lys Gly Arg Thr Ala Cys Cys His Ser Glu Thr Val Val 150 155 160

<sup>&</sup>lt;211> 160

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Homo sapiens

<sup>&</sup>lt;223> Human TGF- alpha

<211> 3776 <212> DNA <213> Mus musculus <220> <221> CDS <222> (91)..(570) <220> <223> Murine TGF-alpha <400> 31 qcqccttttt cccccgcgca caccgcggcg gcgcgcgct actcgccaac cgcagggagc 60 geggtggetg cagcaccetg cgctcggaag atg gtc ccc gcg acc gga cag etc Met Val Pro Ala Thr Gly Gln Leu gct ctg cta gcg ctg ggt atc ctg tta gct gtg tgc cag gct ctg gag 162 Ala Leu Leu Ala Leu Gly Ile Leu Leu Ala Val Cys Gln Ala Leu Glu 15 10 aac agc aca tcc ccc ctg agt gac tca ccc gtg gcg gct gca gtg gtg 210 Asn Ser Thr Ser Pro Leu Ser Asp Ser Pro Val Ala Ala Ala Val Val 30 tet cae tte aac aag tge cea gat tee cae act cag tae tge tte cat Ser His Phe Asn Lys Cys Pro Asp Ser His Thr Gln Tyr Cys Phe His 45 50 gga acc tgc cgg ttt ttg gtg cag gaa gag aag cca gca tgt gtc tgc 306 Gly Thr Cys Arg Phe Leu Val Gln Glu Glu Lys Pro Ala Cys Val Cys 60 65 354 cac tot ggg tac gtg ggt gtt cgc tgt gag cat gca gac ctc ctg gct His Ser Gly Tyr Val Gly Val Arg Cys Glu His Ala Asp Leu Leu Ala 75 80 402 gtg gtg gct gcc agc cag aag caa gcc atc act gcc ctg gtg gtg Val Val Ala Ala Ser Gln Lys Lys Gln Ala Ile Thr Ala Leu Val Val 90 95 100 gtc tcc att gtg gcc ctg gct gtc ctc att atc acc tgt gtg ctg atc 450 Val Ser Ile Val Ala Leu Ala Val Leu Ile Ile Thr Cys Val Leu Ile 105 110 115 cac tgc tgt cag ctc cgc aaa cac tgt gag tgg tgc cgt gcc ctc gtc 498 His Cys Cys Gln Leu Arg Lys His Cys Glu Trp Cys Arg Ala Leu Val tgc aga cat gag aag ccc agc gcc ctc ctg aag gga agg act gct tgc Cys Arg His Glu Lys Pro Ser Ala Leu Leu Lys Gly Arg Thr Ala Cys 600 tgc cac tct gag aca gtg gtc tga agatcccaga ggaggaattt ggccaggtgg Cys His Ser Glu Thr Val Val 160 155 cctatgacag cccaaccaag aaaaggcgtc ttgggacaac acccctggca gtgcccaggc 660

ccatgggaca tgctgggaga ccttccccct cagtgcacaa ctgcctgggc aggcttcttc 720 cttgagagtc ttcaaaactg tgtgataaag ctgcctgctg ggctcgctca gtacacccag 780 agaagaggcc agtggaccac attttaaaga caagttgaac acgaacctca aagggttggc 840 cttcttgcta acccacaccg agaatgaget ggggetgetg teceetgeea gecatgaett 900 ccagactgtt tettetetat gggccateta ceceegtee egagacteea tggttggtgt 960 acaaaatgga caaggggaaa cgtctattgt tctttgaaga caccatggcg tcccatgcgc 1020 cctgacatct cctcaggctg tcgtcaggat gcgtgtctta tttattagat ggataacatg 1080 gttttatttg taatctcttt atgtcaatgt caggcgtccg tgctagcgat gcatgagtgg 1140 cgggccgtga agttacacca aatagaaatg tccagccact ttgataaagt aaggctgcag 1200 ggaggggagc ctccgggggg ccgtagagac agaattcaga tgtgcatcct gctgggaacc 1260 aacctgctat cccaaggctg tgcccctca ggcaagaaag tccctggata catgctgtag 1320 caaacttagc aaggccaatg gcagagagtg tcccccttgt ggggacaaga ggacaaaaga 1380 gcctgtgtgg gcacctagga aggacgcaca ccccaaccct tgattaagtg gattagcgtc 1440 ctcgtgagcc cagctccagc atcagcaatg agctgtgagc caaatccatg aaggggacca 1500 gccctttatt ggcttttgct tatttttaga aggagaaatt tgcctttcct atttatttc 1560 aagcattaag aaaaatagcc cactgttttg ttttgtttcc cctttaataa tgttggcatt 1620 attaacaaaa togotaccaa goototocac cootototgg ggaaggaaat gtgttccctg 1680 agcaccegaa gatgetetga eccattgage ceaccettgt teetgggetg ettgggaetg 1740 tcaggagaca gaatacattc agttgcatat taaagagacc attctccaac tcagaagtga 1800 aaagtotgoo cootoggooc tooagggoot otgaagtaag goatgaggga aggggtactg 1860 ggcataccaa gttggtttga agcttctgtc agaagctgcc tgttcctcat tcttctcctt 1920 tatgaatttt totoototoa tatgatotag goattgatot goocaggtot taaccotttg 1980 ttcacaaggc ctgaagccct gaaagaagaa cctgaaactg ttcagtgctg tcctctaact 2040 gcttagctga cacagctcca ggcagaagtt gtcattcttg ctattatttt ggtttgatgc 2100 atcctcctgc ttcagcaacc tggaagctgg gcttacagat gtgcaccacc acgcccagca 2220 ctttttgatt actagagaaa ataggttaag tttgctttcc ttgggtcata caagtgggtc 2280 aaggccaagt gtcacctgct gctcagacac caatcaacct cactggactt cagccctcta 2340 ttggccccca gtgctctcag tagaccccgc ctgaagtact ggctcactgg agggcaggga 2400 gcaacacaaa tggaggagat gtggggcttg ttttagttta gttcacactg ttagcttctg 2460 atgacaggte ttttetggte tgetggagge teetgggtgt etgaettett ttaaagagtt 2520

tgaactgata tcaaaatatc attgggaact tcctcactta atatcccatc ctgaagatgt 2580 tgtcatccaa gactaagcag aggtggtaaa cgcttagcac gagcactcga tggtgaaggc 2640 agaggatete gggaggacet cagetttagt gteegaacgg teetqttete agqteeaqee 2700 agtogoagoa gocagtacoa toccagtgoo tgtacacaga tgcagotggo tcaaacacot 2760 cccagaagtg ccttcccatc agtctaccct ttgcagagtc agaggctgag actaggttga 2820 gacaagteeg gttggacate gteeetgeee egacgtatga cagecetget gegttggtga 2880 ctttgatttt ggaagtcagt cagacaaacg ctgtgtacct tctcctgtgc catgtcacag 2940 cacagatggg cetggttage tgtaccageg ceeteetgte agttcacact ggeeetetet 3000 tgatatttag tacagctact taatcaggag cactcttagc agctaggcag gtcagagggc 3060 agtacagttg attcagtgac tcaggggact agggccattc ccattaaatg agctgaaatt 3120 aattctagag acattgagac agctcagtta gcaatggcac ttaccaaaat cccagataac 3180 ctgaattcaa tcccctggga tcccatctgg tggaaggaga gaaccggctt gcccaggatg 3240 tettetgatt tetaegtaea egeacaegea cacacaeae egeaegeaea cacaegeaea 3300 cacacgcaca tatgagtaca cacaaagtct aagtttttaa aataaaaatt ctagaaagac 3360 ctctacacct ctagctataa ctttaaacac accatgggac atgttttctc cacggaaaag 3420 cacaattgga gacccagtct gaaatgaaag atagattcca ggtggacata cacagaacca 3480 aaccatgagt tagtttgtca tgtgccccag aaaaaggtgg gaacaataat cccaacatcc 3540 ctttcctctc ccccccaaa aaagggggg aggaggcatg gttggtaccc acccacctgc 3600 actggtgagg gaagtgacta tgtccctgag tcctagatgg tgacgttggt cccttcaagt 3660 ccccaggcca ggtttcctag tgcctatgag gcactggata gaccagcaac actctgacac 3720 tgcctacagg ccccagtcct gtaatgagta ataaacttgt ctctctcca aggcca 3776

```
<210> 32
```

<400> 32

 Met Val
 Pro
 Ala
 Thr
 Gly
 Gln
 Leu
 Ala
 <

<sup>&</sup>lt;211> 159

<sup>&</sup>lt;212> PRT

<sup>&</sup>lt;213> Mus musculus

<sup>&</sup>lt;223> Murine TGF-alpha